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OM protein - protein search, using sw model

Run on:

July 15, 2004, 16:25:44; Search time 55.8955 Seconds

(without alignments)

540.877 Million cell updates/sec

Title:

US-09-423-100-6

Perfect score: 587

Sequence:

1 MFPTIPLSRLFDNAMLRAHR.....IVEQCCTSICSLYQLENYCN 107

Scoring table: BLOSUM62

Gapop 10.0, Gapext 0.5

Searched:

1586107 seqs, 282547505 residues

Total number of hits satisfying chosen parameters: 1586107

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database :

A Geneseq 29Jan04:\*

- 1: geneseqp1980s:\*
- 2: geneseqp1990s:\*
- 3: geneseqp2000s:\*
- 4: geneseqp2001s:\*
- 5: geneseqp2002s:\*
- 6: geneseqp2003as:\*
- 7: geneseqp2003bs:\*
- 8: geneseqp2004s:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

### SUMMARIES

		용				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
 1	. <b>-</b> 587	100.0	107	 2	AAY42860	Aay42860 hGH-mini-
2	555.5	94.6	150	2	AAY42861	Aay42861 Chimeric
3	315.5	53.7	116	2	AAR98897	Aar98897 SOD-proin
4	304	51.8	63	2	AAR68900	Aar68900 Human pro
5	304	51.8	117	2	AAR98896	Aar98896 SOD-proin
6	302.5	51.5	137	2	AAR71692	Aar71692 Mating fa
7	299	50.9	56	2	AAR68901	Aar68901 Human pro
8	299	50.9	56	2	AAR78665	Aar78665 Proinsuli
9	299	50.9	96	2	AAR68899	Aar68899 Human pro

10	299	50.9	96	2	AAR78662	Aar78662	Fusion pr
11	299	50.9	145	2	AAR71694	Aar71694	Mating fa
12	299	50.9	146	2	AAR71695	Aar71695	Mating fa
13	294	50.1	52	2	AAY42859	Aay42859	Human ins
14	293	49.9	57	2	AAR04582	Aar04582	Proinsuli
15	288.5	49.1	160	2	AAR79056	Aar79056	Glycosylp
16	287	48.9	52	2	AAR11899	Aar11899	Example o
17	287	48.9	65	2	AAW47365	Aaw47365	Preproins
18	287	48.9	138	2	AAR87086	Aar87086	pKV142 mo
19	284.5	48.5	58	2	AAR96047	Aar96047	Modified
20	284.5	48.5	59	2	AAR96048	Aar96048	Modified
21	284.5	48.5	65	2	AAR88188	Aar88188	N-termina
22	284.5	48.5	109	1	AAP94645	Aap94645	Amino aci
23	284.5	48.5	123	2	AAW19240	Aaw19240	EEAEPK-MI
24	284.5	48.5	123	2	AAW69160	Aaw69160	DNA const
25	284.5	48.5	124	2	AAW78751	Aaw78751	pAK855 pr
26	284.5	48.5	124	6	ABP55059	Abp55059	Insulin p
27	284.5	48.5	124	6	ABB82578	Abb82578	Synthetic
28	284.5	48.5	125	2	AAW19242	Aaw19242	EEAEPK-MI
29	284	48.4	138	1	AAP94643	Aap94643	Amino aci
30	284	48.4	138	2	AAW04890	Aaw04890	S. cerevi
31	284	48.4	140	2	AAR71693		Mating fa
32	284	48.4	140	2	AAR71690	Aar71690	Mating fa
33	283.5	48.3	53	2	AAR65883	Aar65883	Di-Arg-(B
34	283.5	48.3	53	2	AAW18007	Aaw18007	Insl doub
35	283.5	48.3	117	2	AAW78752	Aaw78752	Protein s
36	283.5	48.3	408	4	AAB30705		A Bacillu
37	282	48.0	94	1	AAP94644	Aap94644	Amino aci
38	282	48.0	120	2	AAW19241		EEAEPK-MI
39	281.5	48.0	60	1	AAP20002	Aap20002	Human pro
40	281.5	48.0	105	1	AAP94648	Aap94648	Amino aci
41	281.5	48.0	153	3	AAY53589	<del>-</del>	Human pre
42	281	47.9	102	1	AAP94649	Aap94649	Signal-le
43	281	47.9	104	2	AAR71684		Yeast sig
44	280.5	47.8	55	1	AAP71019		Sequence
45	280.5	47.8	89	2	AAR88179	Aar88179	Signal pe

## ALIGNMENTS

```
RESULT 1
AAY42860
ID
     AAY42860 standard; protein; 107 AA.
XX
     AAY42860;
AC
XX
DT
     19-JAN-2000 (first entry)
XX
     hGH-mini-proinsulin chimeric protein.
DE
XX
     Insulin; precursor; growth hormone; chaperone; intramolecular; folding;
KW
     conformation; chimeric protein; cleavable; recombinant; production;
KW
KW
     yield.
XX
OS
     Synthetic.
OS
     Homo sapiens.
```

```
XX
PN
    WO9950302-A1.
XX
    07-OCT-1999.
PD
XX
PF
    31-MAR-1998;
                   98WO-CN000052.
XX
                   98WO-CN000052.
PR
    31-MAR-1998;
XX
    (TONG-) TONGHUA GANTECH BIOTECHNOLOGY LTD.
PΑ
XX
    Gan Z;
PΙ
XX
    WPI: 1999-610839/52.
DR
XX
    New chimeric proteins containing human growth hormone fragment, used
PТ
PТ
    particularly for the production of human insulin.
XX
PS
    Claim 13; Page 30; 46pp; English.
XX
CC
    This sequence represents a chimeric protein, hGH-mini-proinsulin. This
CC
    chimeric protein contains an N-terminal fragment of human growth hormone
CC
     (hGH) of the sequence given in AAY42855, a cleavable peptide linker
CC
     (AAY42857), and a human insulin precursor comprising insulin A and B
    chains (AAY42859). The hGH portion of the chimeric protein acts as an
CC
CC
    intramolecular chaperone (IMC) for the insulin precursor, enabling it to
    fold correctly. The cleavable peptide linker has a C-terminal Arg residue
CC
    which enables the hGH portion of the chimeric protein to be removed after
CC
    folding has taken place. Production of recombinant human insulin via an
CC
CC
    hGH-proinsulin chimeric protein can provide human insulin with correctly
    linked cysteine bridges with fewer necessary procedural steps, and hence
CC
    resulting in a higher yield of human insulin. The IMC sequences not only
CC
    protect insulin sequences from intracellular degradation by a
CC
CC
    microorganism host, but also promote the folding of the fused insulin
    precursor, facilitate the solubility of the fusion protein and decrease
CC
CC
    the intermolecular interactions among the fusion proteins, thus allowing
CC
    folding of the fused insulin precursor at commercially useful high
    concentrations. The procedural steps of cyanogen bromide cleavage,
CC
    oxidative sulphitolysis and related purification steps can thus be
CC
    eliminated, along with the use of high concentrations of mercaptan or the
CC
CC
    use of hydrophobic absorbent resins
XX
    Sequence 107 AA;
SO
                         100.0%; Score 587; DB 2; Length 107;
  Query Match
                         100.0%; Pred. No. 2.4e-43;
  Best Local Similarity
  Matches 107; Conservative 0; Mismatches
                                                0;
                                                    Indels
                                                               0; Gaps
                                                                          0;
           1 MFPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNPLGTGPRFVNQH 60
Qy
             Db
           1 MFPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNPLGTGPRFVNQH 60
           61 LCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              Db
           61 LCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
```

```
RESULT 2
AAY42861
    AAY42861 standard; protein; 150 AA.
XX
AC
    AAY42861;
XX
DT
    19-JAN-2000 (first entry)
XX
DE
    Chimeric protein, SEQ ID 7.
XX
KW
    Insulin; precursor; growth hormone; chaperone; intramolecular; folding;
     conformation; chimeric protein; cleavable; recombinant; production;
KW
KW
XX
OS
    Synthetic.
OS
    Homo sapiens.
XX
PN
    WO9950302-A1.
XX
    07-OCT-1999.
PD
XX
PF
    31-MAR-1998;
                    98WO-CN000052.
XX
PR
     31-MAR-1998;
                    98WO-CN000052.
XX
     (TONG-) TONGHUA GANTECH BIOTECHNOLOGY LTD.
PΑ
XX
PΙ
    Gan Z;
XX
    WPI; 1999-610839/52.
DR
XX
PT
    New chimeric proteins containing human growth hormone fragment, used
PT
    particularly for the production of human insulin.
XX
     Claim 14; Page 30-31; 46pp; English.
PS
XX
CC
     This sequence represents a chimeric protein, which contains an N-terminal
CC
     fragment of human growth hormone (hGH) of the sequence given in AAY42856,
CC
     a cleavable peptide linker (AAY42857), and a human insulin precursor
CC
     comprising insulin A and B chains (AAY42859). The hGH portion of the
CC
     chimeric protein acts as an intramolecular chaperone (IMC) for the
CC
     insulin precursor, enabling it to fold correctly. The cleavable peptide
CC
     linker has a C-terminal Arg residue which enables the hGH portion of the
CC
     chimeric protein to be removed after folding has taken place. Production
CC
     of recombinant human insulin via an hGH-proinsulin chimeric protein can
    provide human insulin with correctly linked cysteine bridges with fewer
CC
CC
     necessary procedural steps, and hence resulting in a higher yield of
CC
    human insulin. The IMC sequences not only protect insulin sequences from
CC
     intracellular degradation by a microorganism host, but also promote the
CC
     folding of the fused insulin precursor, facilitate the solubility of the
CC
     fusion protein and decrease the intermolecular interactions among the
CC
     fusion proteins, thus allowing folding of the fused insulin precursor at
CC
     commercially useful high concentrations. The procedural steps of cyanogen
```

bromide cleavage, oxidative sulphitolysis and related purification steps

can thus be eliminated, along with the use of high concentrations of

mercaptan or the use of hydrophobic absorbent resins

CC

CC

CC

XX

```
Query Match
                        94.6%; Score 555.5; DB 2; Length 150;
 Best Local Similarity
                        71.3%; Pred. No. 1.7e-40;
 Matches 107; Conservative
                              0; Mismatches
                                               0;
                                                  Indels
                                                           43; Gaps
                                                                       1;
           1 MFPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP----- 49
Qу
             1 MFPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNPQTSLSFSESIP 60
Db
          50 -----LGTGPRFVNQHLCGSHLVEALYLVCGER 77
Qу
                                          61 TPSNREETQQKSNLELLRISLLLIQSWLEPVQLGTGPRFVNOHLCGSHLVEALYLVCGER 120
Db
          78 GFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             121 GFFYTPKTRGIVEQCCTSICSLYQLENYCN 150
Db
RESULT 3
AAR98897
    AAR98897 standard; protein; 116 AA.
XX
AC
    AAR98897;
XX
DT
    03-FEB-1997 (first entry)
XX
DΕ
    SOD-proinsulin hybrid polypeptide.
XX
KW
    Insulin; proinsulin; hybrid polypeptide; protein folding;
KW
    enzymatic cleavage; cyanogen bromide; sulphitolysis.
XX
OS
    Homo sapiens.,
XX
PN
    W09620724-A1.
XX
PD
    11-JUL-1996.
XX
PF
    29-DEC-1994;
                  94WO-US013268.
XX
PR
    29-DEC-1994;
                  94WO-US013268.
XX
PA
    (BIOT-) BIO-TECHNOLOGY GENERAL CORP.
XX
PΙ
    Hartman JR, Mendelovitz S, Gorecki M;
XX
DR
    WPI; 1996-333766/33.
    N-PSDB; AAT34670.
DR
XX
PT
    Recombinant insulin prodn. by correctly folding pro-insulin hybrid
PT
    polypeptide - then enzymatic cleavage of folded product, does not require
PT
    sulphite protection of SH nor use of cyanogen bromide.
XX
PS
    Example 1B; Fig 7; 69pp; English.
XX
CC
    A new method for the production of recombinant human insulin comprises
CC
    folding a hybrid polypeptide comprising proinsulin under conditions that
```

```
CC
     protein to enzymatic cleavage. The insulin produced can then be purified.
CC
     This sequence is a SOD-insulin B chain-Arg-insulin A chain hybrid
CC
     polypeptide and is encoded by the plasmid construct pDBAST-LAT.
CC
     Transformation of the proper E.coli host cells with pDBAST-LAT results in
     the efficient expression of the proinsulin hybrid polypeptide, useful for
CC
     human insulin production. The method produces recombinant human insulin
CC
CC
     identical to the natural hormone. Hazardous and cumbersome procedures
     involving cyanogen bromide and sulphitolysis to protect SH groups are
CC
CC
     avoided since the entire hybrid polypeptide folds efficiently to the
CC
    native structure even with the leader attached and Cys unprotected
XX
SO
    Sequence 116 AA;
                         53.7%; Score 315.5; DB 2; Length 116;
 Query Match
 Best Local Similarity
                         85.3%; Pred. No. 6.9e-20;
          58; Conservative
                                2; Mismatches
                                                 5; Indels
                                                                           1;
          43 YSFLQNPLGT---GPRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSL 99
Qу
                         Db
          49 HEFGDNTAGSTSAGPRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSL 108
         100 YQLENYCN 107
QУ
             Db
         109 YQLENYCN 116
RESULT 4
AAR68900
    AAR68900 standard; peptide; 63 AA.
XX
AC
    AAR68900;
XX
DT
    25-MAR-2003 (revised)
    02-MAR-1995 (first entry)
XX
DE
    Human pro-insulin 4.
XX
KW
     Pro-insulin; A-chain; B-chain; C-chain; disulphide; mercaptan;
KW
    chaotropic agent.
XX
OS
    Homo sapiens.
XX
ΡN
    EP600372-A1.
XX
PD
    08-JUN-1994.
XX
PF
    25-NOV-1993; 93EP-00118993.
XX
PR
    02-DEC-1992;
                   92DE-04240420.
XX
PΑ
     (FARH ) HOECHST AG.
XX
_{\mathrm{PI}}
    Obermeier R, Gerl M, Ludwig J, Sabel W;
XX
DR
    WPI; 1994-177718/22.
XX
```

permit correct disulphide bond formation and subjecting that folded

CC

```
Prodn. of pro-insulin with correct di:sulphide bridges - by treating
PT
     recombinant precursor protein with mercaptan in alkali and in presence of
PT
     chaotropic agent, then isolation on hydrophobic resin.
XX
PS
     Disclosure; Page 11-12; 15pp; German.
XX
CC
     Pro-insulin is produced by treating recombinant precursor protein with a
CC
     mercaptan to provide 2-10 SH residues per Cys residue, in presence of a
CC
     chaotropic agent and in ag. medium of pH 10-11, treating the prod. with 3
     -50 g hydrophobic adsorber resin per l aq. medium of pH 4-7, isolating
CC
CC
     the adsorbed resin and pro-insulin and desorbing the pro-insulin. This
     method produces pro-insulin with correctly bonded Cys bridges. Compared
CC
CC
     with known methods it involves fewer stages (esp. no sulphitolysis or
CC
     cyanogen bromide cleavage) and overall losses during purification are
     reduced, i.e. the process is quicker and gives better yields. Sequences
CC
     of insulin chain A, B and C are given in AAR68895-97. Sequences of pro-
CC
CC
     insulin 1-4 are given in AAR68898-901. (Updated on 25-MAR-2003 to correct
CC
     PN field.)
XX
SO
     Sequence 63 AA;
  Query Match
                         51.8%; Score 304; DB 2; Length 63;
                                 Pred. No. 3.9e-19;
  Best Local Similarity
                         94.7%;
          54; Conservative
                                0; Mismatches
                                                  3; Indels
                                                                0; Gaps
                                                                           0;
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Qу
                 7 GNSARFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 63
Db
RESULT 5
AAR98896
     AAR98896 standard; protein; 117 AA.
ID
XX
AC
     AAR98896;
XX
DT
     03-FEB-1997 (first entry)
XX
DΕ
     SOD-proinsulin hybrid polypeptide.
XX
KW
     Insulin; proinsulin; hybrid polypeptide; protein folding;
KW
     enzymatic cleavage; cyanogen bromide; sulphitolysis.
XX
OS
     Homo sapiens.
XX
PN
     WO9620724-A1.
XX
PD
     11-JUL-1996.
XX
PF
     29-DEC-1994; 94WO-US013268.
XX
PR
     29-DEC-1994;
                   94WO-US013268.
XX
PΑ
     (BIOT-) BIO-TECHNOLOGY GENERAL CORP.
XX
ΡI
     Hartman JR, Mendelovitz S, Gorecki M;
XX
```

PT

```
WPI; 1996-333766/33.
DR
    N-PSDB; AAT34669.
XX
РΤ
     Recombinant insulin prodn. by correctly folding pro-insulin hybrid
PT
     polypeptide - then enzymatic cleavage of folded product, does not require
PT
     sulphite protection of SH nor use of cyanogen bromide.
XX
PS
    Example 1A; Fig 6; 69pp; English.
XX
CC
    A new method for the production of recombinant human insulin comprises
     folding a hybrid polypeptide comprising proinsulin under conditions that
CC
    permit correct disulphide bond formation and subjecting that folded
CC
CC
    protein to enzymatic cleavage. The insulin produced can then be purified.
CC
    This sequence is a SOD-insulin B chain-Lys-Arg-insulin A chain hybrid
CC
    polypeptide and is encoded by the plasmid construct pBAST-R.
CC
    Transformation of the proper E.coli host cells with pBAST-R results in
CC
    the efficient expression of the proinsulin hybrid polypeptide, useful for
CC
    human insulin production. The method produces recombinant human insulin
    identical to the natural hormone. Hazardous and cumbersome procedures
CC
CC
    involving cyanogen bromide and sulphitolysis to protect SH groups are
CC
    avoided since the entire hybrid polypeptide folds efficiently to the
CC
    native structure even with the leader attached and Cys unprotected
XX
SQ
    Sequence 117 AA;
  Query Match
                         51.8%; Score 304; DB 2; Length 117;
 Best Local Similarity
                         82.6%; Pred. No. 6.8e-19;
          57; Conservative
                                3; Mismatches
                                                 5; Indels
                                                               4; Gaps
                                                                           2;
Qу
          43 YSFLQNPLGT---GPRFVNQHLCGSHLVEALYLVCGERGFFYTPKT-RGIVEQCCTSICS 98
                        : | | |:
Db
          49 HEFGDNTAGSTSAGPRFVNQHLCGSHLIEALYLVCGERGFFYTPKTKRGIVEQCCTSICS 108
          99 LYQLENYCN 107
Qy
             Dh
         109 LYQLENYCN 117
RESULT 6
AAR71692
    AAR71692 standard; protein; 137 AA.
XX
AC
    AAR71692:
XX
DΤ
    25-MAR-2003 (revised)
DT
    20-NOV-1995 (first entry)
XX
DE
    Mating factor alpha 1-Insulin precursor ArgB31.
XX
KW
    Human insulin precursor ArgB31; diabetes; Zinc ion complex;
KW
    mating factor alpha 1.
XX
OS
    Homo sapiens.
XX
FH
                    Location/Qualifiers
    Key
FT
                    1. .85
    Protein
FT
                    /label= mating factor alpha-1
```

DΒ

```
FT
    Peptide
                    86. .116
FT
                    /label= B-chain
FT
    Peptide
                   117. .137
FT
                    /label= A-chain
XX
PN
    WO9507931-A1.
XX
PD
    23-MAR-1995.
XX
PF
    16-SEP-1994;
                  94WO-DK000347.
XX
PR
    17-SEP-1993;
                  93DK-00001044.
PR
    02-FEB-1994;
                  94US-00190829.
XX
    (NOVO ) NOVO-NORDISK AS.
PA
XX
    Havelund S, Halstrom JB, Jonassen I, Andersen AS, Markussen J;
PΙ
XX
DR
    WPI; 1995-131314/17.
    N-PSDB; AAQ86425.
DR
XX
PT
    Acylated insulin deriv. which may be present as a Zinc ion complex - is
PT
    used to treat diabetes and is rapid acting.
XX
PS
    Example 5; Page 78; 100pp; English.
XX
CC
    AAQ86425 encodes AAR71692 mating factor alpha 1-Insulin precursor ArgB31.
CC
    ArgB31 comprises the B and A chains of a claimed human insulin
    derivative. In the final claimed compsn. they are covalently connected
CC
    via disulphide bonds between Cys residues A7/B7 and A20/B19. The
CC
CC
    derivative, which may be present as a zinc ion complex, can be used as a
CC
    fast action treatment for diabetes. (Updated on 25-MAR-2003 to correct PN
    field.)
CC
XX
SO
    Sequence 137 AA;
 Query Match
                        51.5%; Score 302.5; DB 2; Length 137;
                        50.0%; Pred. No. 1.1e-18;
 Best Local Similarity
                              4; Mismatches 27; Indels
           70; Conservative
                                                            39; Gaps
                                                                         4;
           2 FPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQ--KYSFLQ-----N 48
Qу
                    1 : 1
                                    Db
           3 FPSI----FTAVLFAASSALAAPVNTTTEDETAQIPAEAVIGYSDLEGDFDVAVLPFSN 57
Qγ
          49 PLGTG-----PRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 87
                 1
                                      Db
          58 STNNGLLFINTTIASIAAKEEGVSMAKRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 117
          88 IVEQCCTSICSLYQLENYCN 107
Qγ
             118 IVEQCCTSICSLYQLENYCN 137
RESULT 7
AAR68901
TD
    AAR68901 standard; peptide; 56 AA.
XX
```

```
AC
     AAR68901;
XX
DT
     25-MAR-2003
                  (revised)
DT
     02-MAR-1995 (first entry)
XX
DΕ
     Human pro-insulin 3.
XX
KW
     Pro-insulin; A-chain; B-chain; C-chain; disulphide; mercaptan;
KW
     chaotropic agent.
XX
OS
     Homo sapiens.
XX
PN
     EP600372-A1.
XX
     08-JUN-1994.
PD
XX
PF
     25-NOV-1993;
                   93EP-00118993.
XX
PR
     02-DEC-1992;
                   92DE-04240420.
XX
PΑ
     (FARH ) HOECHST AG.
XX
PΙ
     Obermeier R, Gerl M, Ludwig J, Sabel W;
XX
DR
     WPI; 1994-177718/22.
XX
PT
     Prodn. of pro-insulin with correct di:sulphide bridges - by treating
     recombinant precursor protein with mercaptan in alkali and in presence of
PT
PT
     chaotropic agent, then isolation on hydrophobic resin.
XX
PS
     Disclosure; Page 12; 15pp; German.
XX
     Pro-insulin is produced by treating recombinant precursor protein with a
CC
CC
     mercaptan to provide 2-10 SH residues per Cys residue, in presence of a
     chaotropic agent and in aq. medium of pH 10-11, treating the prod. with 3
CC
CC
     -50 g hydrophobic adsorber resin per l aq. medium of pH 4-7, isolating
CC
     the adsorbed resin and pro-insulin and desorbing the pro-insulin. This
     method produces pro-insulin with correctly bonded Cys bridges. Compared
CC
     with known methods it involves fewer stages (esp. no sulphitolysis or
CC
     cyanogen bromide cleavage) and overall losses during purification are
CC
CC
     reduced, i.e. the process is quicker and gives better yields. Sequences
CC
     of insulin chain A, B and C are given in AAR68895-97. Sequences of pro-
     insulin 1-4 are given in AAR68898-901. (Updated on 25-MAR-2003 to correct
CC
CC
     PN field.)
XX
SO
     Sequence 56 AA;
  Query Match
                         50.9%; Score 299; DB 2; Length 56;
  Best Local Similarity
                         100.0%; Pred. No. 9.5e-19;
  Matches
            53; Conservative
                                0; Mismatches
                                                                  Gaps
                                                                            0;
                                                  0; Indels
           55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              4 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 56
Db
```

```
AAR78665
ΙD
     AAR78665 standard; protein; 56 AA.
XX
AC
     AAR78665;
XX
DT
     03-APR-1996 (first entry)
XX
DE
     Proinsulin sequence 3.
XX
KW
     Proinsulin; post-translational modification; recombinant production;
     protein folding; conformation.
KW
XX
OS
     Synthetic.
XX
                     Location/Oualifiers
FH
     Key
FT
     Region
                     1. .4
FT
                     /label= R2
FT
                     /note= "a peptide of 4 amino acids"
FT
     Peptide
                     5. .34
                     /label = R1 - (B2 - B29) - Y
FT
FT
                     /note= "human insulin B-chain"
FT
     Region
                     35
FT
                     /label= X
FT
                     36. .56
     Peptide
FT
                     /label = Gly - (A2 - A20) - R3
                     /note= "human insulin A-chain"
FT
XX
PN
     EP668292-A2.
XX
PD
     23-AUG-1995.
XX
                    95EP-00101748.
PF
     09-FEB-1995;
XX
PR
     18-FEB-1994;
                    94DE-04405179.
XX
PΑ
     (FARH ) HOECHST AG.
XX
PI
     Obermeier R, Gerl M, Ludwig J, Sabel W;
XX
DR
     WPI; 1995-284754/38.
XX
PT
     Isolation of insulin that is correctly post-translationally processed -
     by reacting pro:insulin with a mercaptan in the presence of a chaotropic
PT
     agent and purificn. after absorption to hydrophobic resin.
XX
PS
     Example 2; Page 13; 16pp; German.
XX
CC
     The present sequence is an example of a proinsulin molecule corresp. to
CC
     the general formula R2-R1-(B2-B29)-Y-X-Gly-(A2-A20)-R3 (II). In formula
CC
     (II), X = Lys, Arg or a peptide of 2-35 amino acids contg. Lys or Arg at
CC
     the N- and C-termini; Y = a natural amino acid; R1 = Phe or a bond; R2 =
CC
     H, Arg, Lys, a peptide of 2-45 amino acids contg. Arg or Lys at the N-
CC
     and C-termini; R3 = a natural amino acid; (A2-A20) and (B2-B29) are the
CC
     insulin A- and B-chain sequences from human or other insulin. The
CC
     proinsulin molecule (produced in recombinant E.coli) is reacted with
CC
     mercaptan at a ratio of 2-10 SH residues of mercaptan per Cys residue of
CC
     proinsulin. The reaction takes place in the presence of a chaotropic
```

```
CC
    auxiliary agent at pH 10-11 and results in proinsulin with correctly
    linked cystine bridges. Reaction with trypsin and opt. carboxypeptidase B
CC
CC
    yields correctly folded insulin. The insulin is isolated by absortion on
CC
    a hydrophobic resin
XX
SQ
    Sequence 56 AA;
                         50.9%; Score 299; DB 2; Length 56;
 Query Match
 Best Local Similarity
                        100.0%; Pred. No. 9.5e-19;
          53; Conservative 0; Mismatches
 Matches
                                                0; Indels
                                                               0; Gaps
                                                                           0;
          55 RFVNOHLCGSHLVEALYLVCGERGFFYTPKTRGIVEOCCTSICSLYOLENYCN 107
Qy
             Db
           4 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 56
RESULT 9
AAR68899
ID
    AAR68899 standard; peptide; 96 AA.
XX
    AAR68899;
AC
XX
DT
    25-MAR-2003 (revised)
    02-MAR-1995 (first entry)
DT
XX
    Human pro-insulin 2.
DE
XX
    Pro-insulin; A-chain; B-chain; C-chain; disulphide; mercaptan;
ΚW
KW
    chaotropic agent.
XX
OS
    Homo sapiens.
XX
    EP600372-A1.
PN
XX
    08-JUN-1994.
ΡĎ
XX
    25-NOV-1993;
                 93EP-00118993.
PF
XX
    02-DEC-1992; 92DE-04240420.
PR
XX
    (FARH ) HOECHST AG.
PA
XX
    Obermeier R, Gerl M, Ludwig J, Sabel W;
PΙ
XX
    WPI; 1994-177718/22.
DR
XX
     Prodn. of pro-insulin with correct di:sulphide bridges - by treating
PT
PT
     recombinant precursor protein with mercaptan in alkali and in presence of
PT
     chaotropic agent, then isolation on hydrophobic resin.
XX
PS
     Disclosure; Page 11; 15pp; German.
XX
CC
     Pro-insulin is produced by treating recombinant precursor protein with a
CC
    mercaptan to provide 2-10 SH residues per Cys residue, in presence of a
CC
     chaotropic agent and in aq. medium of pH 10-11, treating the prod. with 3
CC
     -50 g hydrophobic adsorber resin per l aq. medium of pH 4-7, isolating
     the adsorbed resin and pro-insulin and desorbing the pro-insulin. This
CC
```

```
method produces pro-insulin with correctly bonded Cys bridges. Compared
CC
    with known methods it involves fewer stages (esp. no sulphitolysis or
    cyanogen bromide cleavage) and overall losses during purification are
CC
    reduced, i.e. the process is quicker and gives better yields. Sequences
CC
    of insulin chain A, B and C are given in AAR68895-97. Sequences of pro-
CC
CC
    insulin 1-4 are given in AAR68898-901. (Updated on 25-MAR-2003 to correct
CC
    PN field.)
XX
    Sequence 96 AA;
SO
                         50.9%; Score 299; DB 2; Length 96;
 Query Match
                         100.0%; Pred. No. 1.5e-18;
 Best Local Similarity
 Matches 53; Conservative 0; Mismatches
                                                0; Indels
                                                                0; Gaps
                                                                           0;
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             Db
          44 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 96
RESULT 10
AAR78662
    AAR78662 standard; protein; 96 AA.
XX
AC
    AAR78662;
XX
DT
     03-APR-1996 (first entry)
XX
DΕ
     Fusion protein contg. proinsulin sequence 3.
XX
     Proinsulin; post-translational modification; recombinant production;
KW
KW
    protein folding; conformation.
XX
OS
     Synthetic.
XX
                    Location/Qualifiers
FH
     Key
                    41. .44
FT
     Region
                    /label= R2
FT
                    /note= "a peptide of 4 amino acids"
FT
                    45. .74
FT
     Peptide
                    /label = R1 - (B2 - B29) - Y
FT
                    /note= "human insulin B-chain"
FT
FT
                    75
     Region
                    /label= X
FT
                    76. .96
FΤ
     Peptide
FT
                    /label= Gly-(A2-A20)-R3
FT
                    /note= "human insulin A-chain"
XX
ΡN
     EP668292-A2.
XX
     23-AUG-1995.
PD
XX
PF
     09-FEB-1995;
                   95EP-00101748.
XX
PR
     18-FEB-1994;
                   94DE-04405179.
XX
PΑ
     (FARH ) HOECHST AG.
XX
```

CC

```
Obermeier R, Gerl M, Ludwig J, Sabel W;
PI
XX
DR
    WPI; 1995-284754/38.
XX
    Isolation of insulin that is correctly post-translationally processed -
PT
    by reacting pro:insulin with a mercaptan in the presence of a chaotropic
PT
    agent and purificn. after absorption to hydrophobic resin.
PΤ
XX
    Example 2; Page 8; 16pp; German.
PS
XX
    The present sequence is that of a fusion protein, produced in E.coli
CC
    which contains an example of a proinsulin molecule corresp. to the
CC
    general formula R2-R1-(B2-B29)-Y-X-Gly-(A2-A20)-R3 (II). In formula (II),
CC
    X = Lys, Arg or a peptide of 2-35 amino acids contg. Lys or Arg at the N-
CC
    and C-termini; Y = a natural amino acid; R1 = Phe or a bond; R2 = H, Arg,
CC
    Lys, a peptide of 2-45 amino acids contq. Arg or Lys at the N- and C-
CC
CC
     termini; R3 = a natural amino acid; (A2-A20) and (B2-B29) are the insulin
    A- and B-chain sequences from human or other insulin. The proinsulin
CC
    molecule, released by cyanogen bromide, is reacted with mercaptan at a
CC
     ratio of 2-10 SH residues of mercaptan per Cys residue of proinsulin. The
CC
CC
     reaction takes place in the presence of a chaotropic auxiliary agent at
    pH 10-11 and results in proinsulin with correctly linked cystine bridges.
CC
    Reaction with trypsin and opt. carboxypeptidase B yields correctly folded
CC
    insulin. The insulin is isolated by absortion on a hydrophobic resin
CC
XX
    Sequence 96 AA;
SQ
                         50.9%; Score 299; DB 2; Length 96;
  Query Match
                         100.0%; Pred. No. 1.5e-18;
  Best Local Similarity
          53; Conservative 0; Mismatches
                                                    Indels
                                                               0; Gaps
                                                                            0;
  Matches
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qy
             44 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 96
Db
RESULT 11
AAR71694
     AAR71694 standard; protein; 145 AA.
XX
AC
    AAR71694;
XX
DT
     25-MAR-2003 (revised)
     20-NOV-1995 (first entry)
DT
XX
DE
     Mating factor alpha 1-Insulin precursor ArgB1, ArgB31 N-terminal.
XX
KW
     Human insulin precursor ArgB1, ArgB31; diabetes; Zinc ion complex;
     mating factor alpha 1; N-terminal EEAEAEAR.
KW
XX
OS
     Homo sapiens.
XX
                    Location/Qualifiers
FH
     Key
                    1. .85
FT
     Protein
                    /label= mating factor alpha-1
FT
                    86. .93
FT
     Peptide
                    /label= N-terminal peptide
FT
```

```
FT
     Peptide
                    94. .124
                    /label= B-chain
FT
FT
                    125. .145
     Peptide
FT
                    /label= A-chain
XX
PN
    WO9507931-A1.
XX
PD
    23-MAR-1995.
XX
                   94WO-DK000347.
PF
    16-SEP-1994;
XX
                   93DK-00001044.
PR
     17-SEP-1993;
PR
     02-FEB-1994;
                   94US-00190829.
XX
     (NOVO ) NOVO-NORDISK AS.
PΑ
XX
PΙ
    Havelund S, Halstrom JB, Jonassen I, Andersen AS, Markussen J;
XX
DR
    WPI; 1995-131314/17.
    N-PSDB; AAQ86429.
DR
XX
     Acylated insulin deriv. which may be present as a Zinc ion complex - is
PT
PT
     used to treat diabetes and is rapid acting.
XX
PS
     Example 5; Page 82-83; 100pp; English.
XX
CC
     AAQ86429 encodes AAR71694 mating factor alpha 1-Insulin precursor ArgB1,
     ArgB31 N-terminal EEAEAEAR. The insulin precursor comprises the B and A
CC
     chains of a claimed human insulin derivative preceded by the N-terminal
CC
CC
     amino acids EEAEAEAR. In the final claimed compsn. they are covalently
     connected via disulphide bonds between Cys residues A7/B7 and A20/B19.
CC
     The derivative, which may be present as a zinc ion complex, can be used
CC
     as a fast action treatment for diabetes. (Updated on 25-MAR-2003 to
CC
CC
     correct PN field.)
XX
SQ
     Sequence 145 AA;
  Query Match
                         50.9%; Score 299; DB 2; Length 145;
  Best Local Similarity 100.0%; Pred. No. 2.2e-18;
                             0; Mismatches
                                               0; Indels
                                                               0; Gaps
                                                                           0;
  Matches
          53; Conservative
           55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qv
             Db
           93 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 145
RESULT 12
AAR71695
     AAR71695 standard; protein; 146 AA.
ΙD
XX
AC
     AAR71695;
XX
DT
     25-MAR-2003 (revised)
DT
     20-NOV-1995 (first entry)
XX
DE
     Mating factor alpha 1-Insulin precursor ArgB1, ArgB31 N-terminal.
XX
```

```
KW
    Human insulin precursor ArgB1, ArgB31; diabetes; Zinc ion complex;
ΚW
    mating factor alpha 1; N-terminal EEAEAEAER.
XX
OS
    Homo sapiens.
XX
                    Location/Qualifiers
FH
    Key
                    1. .85
FT
    Protein
                    /label= mating factor alpha-1
FT
    Peptide
                    86. .94
FT
FT
                    /label= N-terminal peptide
                    95. .125
FT
    Peptide
                    /label= B-chain
FT
    Peptide
                    126. .146
FT
FT
                    /label= A-chain
XX
PN
    WO9507931-A1.
XX
PD
    23-MAR-1995.
XX
                  94WO-DK000347.
PF
    16-SEP-1994;
XX
PR
    17-SEP-1993;
                   93DK-00001044.
    02-FEB-1994;
                   94US-00190829.
PR
XX
     (NOVO ) NOVO-NORDISK AS.
PA
XX
    Havelund S, Halstrom JB, Jonassen I, Andersen AS, Markussen J;
PΙ
XX
DR
    WPI; 1995-131314/17.
    N-PSDB; AAQ86432.
DR
XX
PT
    Acylated insulin deriv. which may be present as a Zinc ion complex - is
    used to treat diabetes and is rapid acting.
PT
XX
PS
    Example 6; Page 85; 100pp; English.
XX
CC
    AAQ86432 encodes AAR71695 mating factor alpha 1-Insulin precursor ArgB1,
CC
    ArgB31 N-terminal EEAEAEAER. The insulin precursor comprises the B and A
     chains of a claimed human insulin derivative preceded by the N-terminal
CC
     amino acids EEAEAEAER. In the final claimed compsn. they are covalently
CC
CC
     connected via disulphide bonds between Cys residues A7/B7 and A20/B19.
CC
     The derivative, which may be present as a zinc ion complex, can be used
     as a fast action treatment for diabetes. (Updated on 25-MAR-2003 to
CC
CC
     correct PN field.)
XX
SQ
     Sequence 146 AA;
  Query Match
                         50.9%; Score 299; DB 2; Length 146;
  Best Local Similarity 100.0%; Pred. No. 2.3e-18;
  Matches
          53; Conservative 0; Mismatches
                                                 0; Indels
                                                               0; Gaps
                                                                           0;
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              94 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 146
Db
```

```
AAY42859
     AAY42859 standard; protein; 52 AA.
XX
AC
     AAY42859;
XX
DT
     19-JAN-2000 (first entry)
XX
DE
     Human insulin precursor, SEQ ID 5.
XX
     Insulin; precursor; growth hormone; chaperone; intramolecular; folding;
KW
ΚW
     conformation; chimeric protein; cleavable; recombinant; production;
KW
     yield.
XX
OS
     Homo sapiens.
XX
PN
     WO9950302-A1.
XX
PD
     07-OCT-1999.
XX
PF
     31-MAR-1998;
                    98WO-CN000052.
XX
PR
     31-MAR-1998;
                    98WO-CN000052.
XX
PΑ
     (TONG-) TONGHUA GANTECH BIOTECHNOLOGY LTD.
XX
ΡI
     Gan Z;
XX
DR
    WPI; 1999-610839/52.
XX
PT
     New chimeric proteins containing human growth hormone fragment, used
PT
     particularly for the production of human insulin.
XX
PS
     Claim 12; Page 29-30; 46pp; English.
XX
CC
     This sequence represents a human insulin precursor comprising insulin A
CC
     and B chains. This insulin precursor is a component of the chimeric
CC
     proteins hGH-mini-proinsulin (AAY42860) and the chimeric protein given in
CC
     AAY42861. These chimeric proteins additionally contain an N-terminal
CC
     fragment of human growth hormone (hGH) and a cleavable peptide linker
CC
     (AAY42857). The hGH portion of the chimeric protein acts as an
CC
     intramolecular chaperone (IMC) for the insulin precursor, enabling it to
CC
     fold correctly. The cleavable peptide linker has a C-terminal Arg residue
     which enables the hGH portion of the chimeric protein to be removed after
CC
CC
     folding has taken place. Production of recombinant human insulin via an
CC
     hGH-proinsulin chimeric protein can provide human insulin with correctly
```

CC concentrations. The procedural steps of cyanogen bromide cleavage,
CC oxidative sulphitolysis and related purification steps can thus be
CC eliminated, along with the use of high concentrations of mercaptan or the
CC use of hydrophobic absorbent resins
XX

linked cysteine bridges with fewer necessary procedural steps, and hence resulting in a higher yield of human insulin. The IMC sequences not only

microorganism host, but also promote the folding of the fused insulin

folding of the fused insulin precursor at commercially useful high

precursor, facilitate the solubility of the fusion protein and decrease

the intermolecular interactions among the fusion proteins, thus allowing

protect insulin sequences from intracellular degradation by a

CC

CC

CC

CC

CC

CC

```
SQ
    Sequence 52 AA;
 Query Match
                         50.1%; Score 294; DB 2; Length 52;
  Best Local Similarity 100.0%; Pred. No. 2.4e-18;
          52; Conservative 0; Mismatches 0; Indels
 Matches
                                                              0; Gaps
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qy
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 52
RESULT 14
AAR04582
    AAR04582 standard; protein; 57 AA.
ΙD
XX
AC
    AAR04582;
XX
DT
    25-MAR-2003 (revised)
DT
    14-SEP-1990 (first entry)
XX
DΕ
    Proinsulin analogue with a Lys residue linking the A and B chains.
XX
KW
    insulin fusion protein; pro-insulin analogue; tendamistate;
KW
    Lys-Lys bridge; ds.
XX
OS
    Synthetic.
XX
FΗ
                    Location/Qualifiers
    Key
FT
    Peptide
                    1. .35
                    /label= Insulin B chain
FT
FT
    Misc-difference 36
FT
                    /label= Lys residue linking insulin B chain to A chain
FT
                    37. .57
    Peptide
FT
                    /label= Insulin A chain
XX
PN
    EP367163-A.
XX
PD
    09-MAY-1990.
XX
PF
    28-OCT-1989;
                   89EP-00120056.
XX
PR
    03-NOV-1988;
                   88DE-03837273.
    19-AUG-1989;
                   89DE-03927449.
PR
XX
PΑ
     (FARH ) HOECHST AG.
XX
PΙ
    Koller KP, Riess GJ, Uhlmann E, Wallmeier H;
XX
    WPI; 1990-141149/19.
DR
DR
    N-PSDB; AAQ04335.
XX
PT
    New insulin fusion proteins - comprise pro-insulin analogue linked to
PT
    tendamistate.
```

This sequence is joined to the C-terminus of an N-terminal fragment

XX PS

XX CC Disclosure; Page ?; -pp; German.

0;

```
CC
     comprising opt. modified tendamistate. This fusion protein may be
CC
     converted into human insulin using known methods. The synthetic gene was
CC
    prepared by the phosphoramidite method. See also AAQ04336. (Updated on 25
CC
    -MAR-2003 to correct PR field.) (Updated on 25-MAR-2003 to correct PI
CC
    field.)
XX
SQ
    Sequence 57 AA;
 Query Match
                         49.9%; Score 293; DB 2; Length 57;
 Best Local Similarity 96.2%; Pred. No. 3.2e-18;
 Matches
          51; Conservative
                                2; Mismatches 0; Indels
                                                               0; Gaps
                                                                           0;
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
QУ
             5 KFVNQHLCGSHLVEALYLVCGERGFFYTPKTKGIVEQCCTSICSLYQLENYCN 57
Db
RESULT 15
AAR79056
    AAR79056 standard; protein; 160 AA.
XX
AC
    AAR79056;
XX
DT
    25-MAR-2003 (revised)
DT
    24-JAN-1996 (first entry)
XX
DE
    Glycosylphosphatidylinositol-anchored human recombinant insulin.
XX
KW
    GPI; glycosylphophatidylinositol; insulin; hormone; solubilization;
KW
    Saccharomyces cerevisiae; anchor; Gas1; plasmid pBY40.
XX
OS
    Homo sapiens.
XX
FH
                    Location/Qualifiers
FT
    Misc-difference 44. .129
FT
                    /note= "anchor attachment site"
XX
PN
    WO9522614-A1.
XX
PD
    24-AUG-1995.
XX
PF
    16-FEB-1995;
                   95WO-BR000010.
XX
PR
    17-FEB-1994;
                   94BR-00000600.
XX
PA
     (FINE-) FINEP FINANCIADORA ESTUDOS & PROJETOS.
     (ESCO-) ESCOLA PAULISTA MEDICINA.
PA
XX
PI
    Cardoso De Almeida ML, Amaral De Castilho Valavicius;
PΙ
    Gomes De Amorim Filho A;
XX
DR
    WPI; 1995-302720/39.
DR
    N-PSDB; AAQ99460.
XX
PT
    Recombinant prodn. of proteins, e.g. insulin - by producing the protein
PT
    with a glycosyl:phosphatidyl:inositol anchor followed by selective
PT
    release.
```

```
XX
PS
    Disclosure; Fig 3; 51pp; English.
XX
CC
    Human recombinant insulin may be expressed in Saccharomyces cerevisiae
CC
    following linkage of the gene to the glycosylphospatidylinositol anchor.
CC
    This anchoring technique can provide for the release of the product in a
CC
    highly specific and selective manner. In addition, the recombinant
CC
    protein will contain an epitope which can be used in its final
CC
    purification by immunoaffinity. The protein product can be released by
    e.g. nitrous deamination or treatment with neutral detergent. (Updated on
CC
CC
    25-MAR-2003 to correct PI field.)
XX
SQ
    Sequence 160 AA;
 Query Match
                        49.1%; Score 288.5; DB 2; Length 160;
 Best Local Similarity
                        98.1%;
                               Pred. No. 2e-17;
 Matches 53; Conservative
                              0; Mismatches
                                              0;
                                                   Indels
                                                             1; Gaps
                                                                        1;
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKT-RGIVEQCCTSICSLYQLENYCN 107
Qу
             Db
          43 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTKRGIVEQCCTSICSLYQLENYCN 96
```

Search completed: July 15, 2004, 16:35:35 Job time: 56.8955 secs

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 15, 2004, 16:30:45; Search time 16.1698 Seconds

(without alignments)

341.624 Million cell updates/sec

Title: US-09-423-100-6

Perfect score: 587

Sequence: 1 MFPTIPLSRLFDNAMLRAHR.....IVEQCCTSICSLYQLENYCN 107

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 389414 seqs, 51625971 residues

Total number of hits satisfying chosen parameters: 389414

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database : Issued Patents AA:\*

1: /cgn2 6/ptodata/2/iaa/5A COMB.pep:\*

2: /cgn2\_6/ptodata/2/iaa/5B\_COMB.pep:\*

3: /cgn2\_6/ptodata/2/iaa/6A\_COMB.pep:\*

4: /cgn2\_6/ptodata/2/iaa/6B\_COMB.pep:\*

5: /cgn2\_6/ptodata/2/iaa/PCTUS\_COMB.pep:\*

6: /cgn2 6/ptodata/2/iaa/backfiles1.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

			ક				
R	esult		Query				
	No.	Score	Match	Length	DB	ID	Description
	1	304	51.8	63	1	US-08-160-376A-6	Sequence 6, Appli
	2	302.5	51.5	137	1	US-08-400-256-39	Sequence 39, Appl
	3	302.5	51.5	137	3	US-08-975-365-39	Sequence 39, Appl
	4	299.5	51.0	66	1	US-08-291-060B-5	Sequence 5, Appli
	5	299	50.9	56	1	US-08-160-376A-7	Sequence 7, Appli
	6	299	50.9	56	1	US-08-389-487-11	Sequence 11, Appl
	7	299	50.9	96	1	US-08-160-376A-5	Sequence 5, Appli
	8	299	50.9	96	1	US-08-389-487-8	Sequence 8, Appli
	9	299	50.9	145	1	US-08-400-256-45	Sequence 45, Appl
	10	299	50.9	145	3	US-08-975-365-45	Sequence 45, Appl
	11	299	50.9	146	1	US-08-400-256-48	Sequence 48, Appl

12	299	50.9	146	3	US-08-975-365-48	Sequence	48. Appl
13	293	49.9	57	1	US-08-030-731A-44	Sequence	
14	287	48.9	65	3	US-08-900-574-3	Sequence	
15	286.5	48.8	66	3	US-08-900-574-5	Sequence	
16	286	48.7	67	3	US-08-900-574-7	Sequence	
17	284.5	48.5	65	1	US-08-468-674B-71	Sequence	
18	284.5	48.5	65	1	US-08-780-571-71	Sequence	
19	284.5	48.5	124	3	US-09-012-669F-36	Sequence	
20	284	48.4	138	3	US-08-932-082-19	Sequence	
21	284	48.4	138	4	US-09-861-687-19	Sequence	
22	284	48.4	140	1	US-08-400-256-33	Sequence	
23	284	48.4	140	1	US-08-400-256-42	Sequence	
24	284	48.4	140	3	US-08-975-365-33	Sequence	
25	284	48.4	140	3	US-08-975-365-42	Sequence	
26	283.5	48.3	53	1	US-08-233-617-4	Sequence	
27	283.5	48.3	53	4	US-08-981-988A-42	Sequence	
28	283.5	48.3	117	3	US-09-012-669F-37	Sequence	
29	281	47.9	104	1	US-08-400-256-15	Sequence	
30	281	47.9	104	3	US-08-975-365-15	Sequence	
31	280.5	47.8	89	1	US-08-468-674B-41	Sequence	41, Appl
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33	280.5	47.8	91	1	US-08-468-674B-45	Sequence	
34	280.5	47.8	91	1	US-08-780-571-45	Sequence	45, Appl
35	280.5	47.8	124	1	US-08-446-646-3	Sequence	3, Appli
36	279.5	47.6	167	1	US-07-918-953-8	Sequence	8, Appli
37	279.5	47.6	167	1	US-08-081-661-8	Sequence	8, Appli
38	278.5	47.4	51	4	US-09-477-924-3	Sequence	3, Appli
39	278.5	47.4	51	4	US-09-723-981-3	Sequence	3, Appli
40	278.5	47.4	51	4	US-09-723-896-3	Sequence	3, Appli
41	278	47.4	117	4	US-09-280-030-63	Sequence	
42	277.5	47.3	53	1	US-08-233-617-3	Sequence	3, Appli
43	277	47.2	96	2	US-09-134-836-4	Sequence	4, Appli
44	277	47.2	96	4	US-09-386-303A-4	Sequence	4, Appli
45	277	47.2	97	1	US-08-160-376A-4	Sequence	4, Appli

### ALIGNMENTS

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RESULT 1
US-08-160-376A-6
; Sequence 6, Application US/08160376A
; Patent No. 5473049
  GENERAL INFORMATION:
    APPLICANT: Obermeier, Ranier
    APPLICANT: Gerl, Martin
    APPLICANT: Ludwig, Jurgen
    APPLICANT: Sabel, Walter
    TITLE OF INVENTION: Process For Obtaining Proinsulin
    TITLE OF INVENTION: Possessing Correctly Linked
    TITLE OF INVENTION: Cystine Bridges
    NUMBER OF SEQUENCES:
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Kenneth A. Genoni, Esq.
      STREET: Rt. 202-206 No. 5473049th/P.O. Box 2500
      CITY: Somerville
      STATE: New Jersey
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COUNTRY: U.S.A.
      ZIP: 08876-1258
    COMPUTER READABLE FORM:
      MEDIUM TYPE: DISKETTE, 3.5 INCH, 1.44 Mb STORAGE
      COMPUTER: IBM 386
      OPERATING SYSTEM: WINDOWS 3.1
      SOFTWARE: WORDPERFECT 5.1
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/160,376A
      FILING DATE: December 1, 1993
      CLASSIFICATION: 530
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: GE P 4240420.7
      FILING DATE: December 2, 1992
    ATTORNEY/AGENT INFORMATION:
      NAME: Barbara V. Maurer, Esq.
      REGISTRATION NUMBER: 31,287
      REFERENCE/DOCKET NUMBER: HOE 92/F 384
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (908) 231-4079
      TELEFAX: (908) 231-2255
  INFORMATION FOR SEQ ID NO: 6:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 63 Amino Acids
      TYPE: Amino Acid (AA)
      TOPOLOGY: not relevant
US-08-160-376A-6
 Query Match
                        51.8%; Score 304; DB 1; Length 63;
 Best Local Similarity 94.7%; Pred. No. 8.3e-29;
          54; Conservative
                               0; Mismatches 3; Indels
 Matches
                                                             0; Gaps
                                                                         0;
          51 GTGPRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qy
                Db
           7 GNSARFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 63
RESULT 2
US-08-400-256-39
; Sequence 39, Application US/08400256
; Patent No. 5750497
  GENERAL INFORMATION:
    APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
                Jonassen, Ib
    APPLICANT:
    APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: No. 57504970 No. 5750497disk of No. 5750497th America, Inc.
      STREET: 405 Lexington Avenue, 64th Floor
      CITY: New York
      STATE: New York
      COUNTRY: United States of America
      ZIP: 10174-6401
    COMPUTER READABLE FORM:
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MEDIUM TYPE: Floppy disk
     COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
   CURRENT APPLICATION DATA:
     APPLICATION NUMBER: US/08/400,256
     FILING DATE: 03-MAR-1995
     CLASSIFICATION: 514
   ATTORNEY/AGENT INFORMATION:
    NAME: Lambiris, Elias J.
     REGISTRATION NUMBER: 33,728
     REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 39:
    SEQUENCE CHARACTERISTICS:
     LENGTH: 137 amino acids
      TYPE: amino acid
     TOPOLOGY: linear
   MOLECULE TYPE: protein
US-08-400-256-39
 Query Match
                     51.5%; Score 302.5; DB 1; Length 137;
 Best Local Similarity 50.0%; Pred. No. 3.3e-28;
 Matches 70; Conservative 4; Mismatches 27; Indels 39; Gaps 4;
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           3 FPSI----FTAVLFAASSALAAPVNTTTEDETAQIPAEAVIGYSDLEGDFDVAVLPFSN 57
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     49 PLGTG-----PRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 87
Qу
           58 STNNGLLFINTTIASIAAKEEGVSMAKRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 117
Db
Qу
        88 IVEQCCTSICSLYQLENYCN 107
           118 IVEQCCTSICSLYQLENYCN 137
RESULT 3
US-08-975-365-39
; Sequence 39, Application US/08975365
; Patent No. 6011007
; GENERAL INFORMATION:
   APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
   APPLICANT: Jonassen, Ib
    APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
   CORRESPONDENCE ADDRESS:
     ADDRESSEE: No. 6011007o No. 6011007disk of No. 6011007th America, Inc.
     STREET: 405 Lexington Avenue, 64th Floor
     CITY: New York
     STATE: New York
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COUNTRY: United States of America
      ZIP: 10174-6401
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/975,365
     FILING DATE:
      CLASSIFICATION: 514
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 08/400,256
      FILING DATE: 03-MAR-1995
    ATTORNEY/AGENT INFORMATION:
    NAME: Lambiris, Elias J.
      REGISTRATION NUMBER: 33,728
      REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 39:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 137 amino acids
      TYPE: amino acid
      TOPOLOGY: linear
    MOLECULE TYPE: protein
US-08-975-365-39
                       51.5%; Score 302.5; DB 3; Length 137;
 Query Match
 Best Local Similarity 50.0%; Pred. No. 3.3e-28;
 Matches 70; Conservative 4; Mismatches 27; Indels 39; Gaps
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Qу
            3 FPSI----FTAVLFAASSALAAPVNTTTEDETAQIPAEAVIGYSDLEGDFDVAVLPFSN 57
Db
         49 PLGTG-----PRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 87
QУ
                                    - 1
         58 STNNGLLFINTTIASIAAKEEGVSMAKRFVNQHLCGSHLVEALYLVCGERGFFYTPKTRG 117
Db
         88 IVEQCCTSICSLYQLENYCN 107
Qу
            118 IVEQCCTSICSLYQLENYCN 137
Db
RESULT 4
US-08-291-060B-5
; Sequence 5, Application US/08291060B
; Patent No. 5728543
  GENERAL INFORMATION:
    APPLICANT: Dorschug, Michael
    APPLICANT: Koller, Klaus-Peter
    APPLICANT: Marquardt, Rudiger
    APPLICANT: Meiwes, Johannes
    TITLE OF INVENTION: An Enzymatic Process for the
    TITLE OF INVENTION: Conversion of Preproinsulins Into Insulins
```

```
;
    NUMBER OF SEQUENCES: 5
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Finnegan, Henderson, Farabow, Garrett &
      ADDRESSEE: Dunner, L.L.P.
      STREET: 1300 I Street, N.W.
      CITY: Washington
      STATE: D.C.
      COUNTRY: USA
      ZIP: 20005-3315
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.30
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/291,060B
      FILING DATE: 08-AUG-1994
      CLASSIFICATION: 435
    ATTORNEY/AGENT INFORMATION:
      NAME: Einaudi, Carol P.
      REGISTRATION NUMBER: 32,220
      REFERENCE/DOCKET NUMBER: 02481.1105-02000
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (202) 408-4366
      TELEFAX: (202) 408-4400
  INFORMATION FOR SEQ ID NO: 5:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 66 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
US-08-291-060B-5
                        51.0%; Score 299.5; DB 1; Length 66;
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 Matches 55; Conservative
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Qу
             8 DPNSNG-RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 66
Db
RESULT 5
US-08-160-376A-7
; Sequence 7, Application US/08160376A
; Patent No. 5473049
  GENERAL INFORMATION:
    APPLICANT: Obermeier, Ranier
    APPLICANT: Gerl, Martin
    APPLICANT: Ludwig, Jurgen
    APPLICANT: Sabel, Walter
    TITLE OF INVENTION: Process For Obtaining Proinsulin
    TITLE OF INVENTION: Possessing Correctly Linked
    TITLE OF INVENTION: Cystine Bridges
    NUMBER OF SEQUENCES: 7
    CORRESPONDENCE ADDRESS:
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STREET: Rt. 202-206 No. 5473049th/P.O. Box 2500
      CITY: Somerville
      STATE: New Jersey
      COUNTRY: U.S.A.
      ZIP: 08876-1258
    COMPUTER READABLE FORM:
      MEDIUM TYPE: DISKETTE, 3.5 INCH, 1.44 Mb STORAGE
      COMPUTER: IBM 386
      OPERATING SYSTEM: WINDOWS 3.1
      SOFTWARE: WORDPERFECT 5.1
   CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/160,376A
      FILING DATE: December 1, 1993
      CLASSIFICATION: 530
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: GE P 4240420.7
      FILING DATE: December 2, 1992
    ATTORNEY/AGENT INFORMATION:
      NAME: Barbara V. Maurer, Esq.
      REGISTRATION NUMBER: 31,287
      REFERENCE/DOCKET NUMBER: HOE 92/F 384
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (908) 231-4079
      TELEFAX: (908) 231-2255
  INFORMATION FOR SEQ ID NO: 7:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 56 Amino Acids
      TYPE: Amino Acid (AA)
      TOPOLOGY: not relevant
US-08-160-376A-7
                        50.9%; Score 299; DB 1; Length 56;
 Query Match
 Best Local Similarity 100.0%; Pred. No. 2.8e-28;
                              0; Mismatches
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                                                              0; Gaps
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 Matches 53; Conservative
          55 RFVNOHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Οv
             4 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 56
RESULT 6
US-08-389-487-11
; Sequence 11, Application US/08389487
; Patent No. 5663291
  GENERAL INFORMATION:
    APPLICANT: Obermeier, Rainer
    APPLICANT: Gerl, Martin
    APPLICANT: Ludwig, Jurgen
    APPLICANT: Sabel, Walter
    TITLE OF INVENTION: Process for Obtaining Insulin Having
    TITLE OF INVENTION: Correctly Linked Cystine Bridges
    NUMBER OF SEQUENCES: 12
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Finnegan, Henderson, Farabow, Garrett &
      ADDRESSEE: Dunner
      STREET: 1300 I Street, N.W.
```

ADDRESSEE: Kenneth A. Genoni, Esq.

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CITY: Washington
       STATE: D.C.
      COUNTRY: United States of America
       ZIP: 20005-3315
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
    CURRENT APPLICATION DATA:
     APPLICATION NUMBER: US/08/389,487
      FILING DATE:
      CLASSIFICATION: 530
    ATTORNEY/AGENT INFORMATION:
     NAME: Einaudi, Carol P.
      REGISTRATION NUMBER: 32,220
      REFERENCE/DOCKET NUMBER: 02481.1424-00000
    TELECOMMUNICATION INFORMATION:
       TELEPHONE: 202-408-4000
       TELEFAX: 202-408-4400
   INFORMATION FOR SEQ ID NO: 11:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 56 amino acids
      TYPE: amino acid
       STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
US-08-389-487-11
  Query Match 50.9%; Score 299; DB 1; Length 56; Best Local Similarity 100.0%; Pred. No. 2.8e-28;
 Matches 53; Conservative 0; Mismatches 0; Indels
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                                                                               0;
           55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              4 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 56
Db
RESULT 7
US-08-160-376A-5
; Sequence 5, Application US/08160376A
; Patent No. 5473049
; GENERAL INFORMATION:
     APPLICANT: Obermeier, Ranier
    APPLICANT: Gerl, Martin
APPLICANT: Ludwig, Jurgen
APPLICANT: Sabel, Walter
   TITLE OF INVENTION: Process For Obtaining Proinsulin TITLE OF INVENTION: Possessing Correctly Linked
     TITLE OF INVENTION: Cystine Bridges
    NUMBER OF SEQUENCES: 7
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Kenneth A. Genoni, Esq.
      STREET: Rt. 202-206 No. 5473049th/P.O. Box 2500
   CITY: Somerville
     STATE: New Jersey
     COUNTRY: U.S.A.
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ZIP: 08876-1258
    COMPUTER READABLE FORM:
      MEDIUM TYPE: DISKETTE, 3.5 INCH, 1.44 Mb STORAGE
      COMPUTER: IBM 386
      OPERATING SYSTEM: WINDOWS 3.1
      SOFTWARE: WORDPERFECT 5.1
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/160,376A
      FILING DATE: December 1, 1993
      CLASSIFICATION: 530
    PRIOR APPLICATION DATA:
;
      APPLICATION NUMBER: GE P 4240420.7
      FILING DATE: December 2, 1992
    ATTORNEY/AGENT INFORMATION:
      NAME: Barbara V. Maurer, Esq.
      REGISTRATION NUMBER: 31,287
      REFERENCE/DOCKET NUMBER: HOE 92/F 384
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (908) 231-4079
      TELEFAX: (908) 231-2255
  INFORMATION FOR SEQ ID NO: 5:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 96 Amino Acids
      TYPE: Amino Acid (AA)
      TOPOLOGY: not relevant
US-08-160-376A-5
 Query Match
                        50.9%; Score 299; DB 1; Length 96;
 Best Local Similarity 100.0%; Pred. No. 5.5e-28;
 Matches 53; Conservative 0; Mismatches 0; Indels
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          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             44 RFVNOHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 96
Db
RESULT 8
US-08-389-487-8
; Sequence 8, Application US/08389487
; Patent No. 5663291
  GENERAL INFORMATION:
    APPLICANT: Obermeier, Rainer
    APPLICANT: Gerl, Martin
    APPLICANT: Ludwig, Jurgen
    APPLICANT: Sabel, Walter
    TITLE OF INVENTION: Process for Obtaining Insulin Having
    TITLE OF INVENTION: Correctly Linked Cystine Bridges
    NUMBER OF SEQUENCES: 12
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Finnegan, Henderson, Farabow, Garrett &
      ADDRESSEE: Dunner
      STREET: 1300 I Street, N.W.
      CITY: Washington
      STATE: D.C.
      COUNTRY: United States of America
      ZIP: 20005-3315
    COMPUTER READABLE FORM:
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MEDIUM TYPE: Floppy disk
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      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
;
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/389,487
      FILING DATE:
      CLASSIFICATION: 530
    ATTORNEY/AGENT INFORMATION:
      NAME: Einaudi, Carol P.
      REGISTRATION NUMBER: 32,220
      REFERENCE/DOCKET NUMBER: 02481.1424-00000
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 202-408-4000
      TELEFAX: 202-408-4400
  INFORMATION FOR SEQ ID NO: 8:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 96 amino acids
      TYPE: amino acid
      STRANDEDNESS: single
      TOPOLOGY: linear
    MOLECULE TYPE: peptide
US-08-389-487-8
                        50.9%; Score 299; DB 1; Length 96;
 Query Match
 Best Local Similarity 100.0%; Pred. No. 5.5e-28;
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         53; Conservative 0; Mismatches
                                               0; Indels
                                                              0; Gaps
          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             44 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 96
Db
RESULT 9
US-08-400-256-45
; Sequence 45, Application US/08400256
; Patent No. 5750497
  GENERAL INFORMATION:
    APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
    APPLICANT: Jonassen, Ib
    APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: No. 57504970 No. 5750497disk of No. 5750497th America, Inc.
      STREET: 405 Lexington Avenue, 64th Floor
      CITY: New York
      STATE: New York
      COUNTRY: United States of America
      ZIP: 10174-6401
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
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CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/400,256
      FILING DATE: 03-MAR-1995
      CLASSIFICATION: 514
    ATTORNEY/AGENT INFORMATION:
     NAME: Lambiris, Elias J.
      REGISTRATION NUMBER: 33,728
      REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 45:
    SEQUENCE CHARACTERISTICS:
     LENGTH: 145 amino acids
      TYPE: amino acid
      TOPOLOGY: linear
    MOLECULE TYPE: protein
US-08-400-256-45
                         50.9%; Score 299; DB 1; Length 145;
  Query Match
  Best Local Similarity 100.0%; Pred. No. 9.1e-28;
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          55 RFVNOHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              93 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 145
Db
RESULT 10
US-08-975-365-45
; Sequence 45, Application US/08975365
: Patent No. 6011007
  GENERAL INFORMATION:
    APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
APPLICANT: Jonassen, Ib
APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: No. 60110070 No. 6011007disk of No. 6011007th America, Inc.
      STREET: 405 Lexington Avenue, 64th Floor
      CITY: New York
       STATE: New York
       COUNTRY: United States of America
       ZIP: 10174-6401
     COMPUTER READABLE FORM:
       MEDIUM TYPE: Floppy disk
       COMPUTER: IBM PC compatible
       OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
;
    CURRENT APPLICATION DATA:
       APPLICATION NUMBER: US/08/975,365
       FILING DATE:
       CLASSIFICATION: 514
     PRIOR APPLICATION DATA:
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APPLICATION NUMBER: US 08/400,256
      FILING DATE: 03-MAR-1995
    ATTORNEY/AGENT INFORMATION:
      NAME: Lambiris, Elias J.
      REGISTRATION NUMBER: 33,728
      REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 45:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 145 amino acids
      TYPE: amino acid
      TOPOLOGY: linear
    MOLECULE TYPE: protein
US-08-975-365-45
 Query Match
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 Best Local Similarity
 Matches 53; Conservative 0; Mismatches
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Qу
             Db
          93 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 145
RESULT 11
US-08-400-256-48
; Sequence 48, Application US/08400256
; Patent No. 5750497
  GENERAL INFORMATION:
    APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
    APPLICANT: Jonassen, Ib
    APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
;
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: No. 57504970 No. 5750497disk of No. 5750497th America, Inc.
      STREET: 405 Lexington Avenue, 64th Floor
      CITY: New York
      STATE: New York
      COUNTRY: United States of America
      ZIP: 10174-6401
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/400,256
      FILING DATE: 03-MAR-1995
      CLASSIFICATION: 514
    ATTORNEY/AGENT INFORMATION:
      NAME: Lambiris, Elias J.
      REGISTRATION NUMBER: 33,728
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REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 48:
   SEQUENCE CHARACTERISTICS:
     LENGTH: 146 amino acids
      TYPE: amino acid
      TOPOLOGY: linear
    MOLECULE TYPE: protein
US-08-400-256-48
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  Query Match
  Best Local Similarity 100.0%; Pred. No. 9.2e-28;
         53; Conservative 0; Mismatches 0; Indels
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          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             94 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 146
RESULT 12
US-08-975-365-48
; Sequence 48, Application US/08975365
; Patent No. 6011007
  GENERAL INFORMATION:
    APPLICANT: Havelund, Svend
    APPLICANT: Halstrom, John
    APPLICANT: Jonassen, Ib
    APPLICANT: Andersen, Asser Sloth
    APPLICANT: Markussen, Jan
    TITLE OF INVENTION: ACYLATED INSULIN
    NUMBER OF SEQUENCES: 49
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: No. 60110070 No. 6011007disk of No. 6011007th America, Inc.
      STREET: 405 Lexington Avenue, 64th Floor
      CITY: New York
      STATE: New York
      COUNTRY: United States of America
      ZIP: 10174-6401
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25
    CURRENT APPLICATION DATA:
    APPLICATION NUMBER: US/08/975,365
      FILING DATE:
      CLASSIFICATION: 514
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 08/400,256
      FILING DATE: 03-MAR-1995
    ATTORNEY/AGENT INFORMATION:
      NAME: Lambiris, Elias J.
      REGISTRATION NUMBER: 33,728
      REFERENCE/DOCKET NUMBER: 3985.220-US
    TELECOMMUNICATION INFORMATION:
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TELEPHONE: 212-867-0123
      TELEFAX: 212-878-9655
  INFORMATION FOR SEQ ID NO: 48:
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      LENGTH: 146 amino acids
      TYPE: amino acid
      TOPOLOGY: linear
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US-08-975-365-48
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Qу
             94 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 146
Db
RESULT 13
US-08-030-731A-44
; Sequence 44, Application US/08030731A
; Patent No. 5426036
  GENERAL INFORMATION:
    APPLICANT: Koller, Klaus-Peter
    APPLICANT: Riess, Guenther Johannes
    APPLICANT: Uhlmann, Eugen
APPLICANT: Wallmeier, Holger
    TITLE OF INVENTION: Processes for the Preparation of Foreign
    TITLE OF INVENTION: Proteins in Streptomycetes
    NUMBER OF SEQUENCES: 48
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Finnegan, Henderson, Farabow, Garrett &
      ADDRESSEE: Dunner
      STREET: 1300 I Street, N.W., Suite 700
      CITY: Washington
      STATE: D.C.
      COUNTRY: USA
      ZIP: 20005-3315
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
       SOFTWARE: PatentIn Release #1.0, Version #1.25
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/030,731A
      FILING DATE: 12-MAR-1993
       CLASSIFICATION: 435
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 07/189,840
       FILING DATE: 03-MAY-1988
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 07/430,622
      FILING DATE: 01-NOV-1989
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: US 07/687,610
      FILING DATE: 19-APR-1991
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PRIOR APPLICATION DATA:
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      FILING DATE: 29-JUL-1991
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;
      FILING DATE: 05-MAY-1987
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: DE P 38 37 273.8
      FILING DATE: 03-NOV-1988
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: DE P 39 27 449.7
      FILING DATE: 19-AUG-1989
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    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: DE P 40 12 818.0
      FILING DATE: 21-APR-1990
    ATTORNEY/AGENT INFORMATION:
      NAME: Kirschner Michael K.
      REGISTRATION NUMBER: 34,851
      REFERENCE/DOCKET NUMBER: 02481-0593-02000
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: 202-408-4000
      TELEFAX: 202-408-4400
   INFORMATION FOR SEQ ID NO:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 57 amino acids
      TYPE: amino acid
      TOPOLOGY: unknown
    MOLECULE TYPE: peptide
US-08-030-731A-44
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                         49.9%; Score 293; DB 1; Length 57;
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          55 RFVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             5 KFVNQHLCGSHLVEALYLVCGERGFFYTPKTKGIVEQCCTSICSLYQLENYCN 57
RESULT 14
US-08-900-574-3
; Sequence 3, Application US/08900574
; Patent No. 6221837
  GENERAL INFORMATION:
    APPLICANT: Ertl, Johann
    APPLICANT: Habermann, Paul
    APPLICANT: Geisen, Karl
                Seipke, Gerhard
    APPLICANT:
     TITLE OF INVENTION: Insulin derivatives with increased zinc
     TITLE OF INVENTION: binding
    NUMBER OF SEQUENCES: 18
    CORRESPONDENCE ADDRESS:
      ADDRESSEE: Finnegan, Henderson, Farabow, Garrett,
      ADDRESSEE: & Dunner, L.L.P.
      STREET: 1300 I Street, N.W.
      CITY: Washington
      STATE: District of Columbia
```

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COUNTRY: U.S.A.
      ZIP: 20005-3315
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25 (EPO)
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/900,574
      FILING DATE: July 24, 1997
      CLASSIFICATION: 514
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: German Application No. 6221837 19630242.0
      FILING DATE: July 26, 1996
    ATTORNEY/AGENT INFORMATION:
      NAME: Carol P. Einaudi
      REGISTRATION NUMBER: 32,220
      REFERENCE/DOCKET NUMBER: 02481.1499-00000
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (202) 408-4000
      TELEFAX: (202) 408-4400
  INFORMATION FOR SEQ ID NO: 3:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 65 amino acids
      TYPE: Amino acid
      STRANDEDNESS: Single
      TOPOLOGY: linear
    MOLECULE TYPE: Protein
    ORIGINAL SOURCE:
      ORGANISM: Escherichia coli
    FEATURE:
      NAME/KEY: Protein
      LOCATION: 1..65
US-08-900-574-3
 Query Match
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RESULT 15
US-08-900-574-5
; Sequence 5, Application US/08900574
; Patent No. 6221837
  GENERAL INFORMATION:
    APPLICANT: Ertl, Johann
    APPLICANT: Habermann, Paul
    APPLICANT: Geisen, Karl
    APPLICANT: Seipke, Gerhard
    TITLE OF INVENTION: Insulin derivatives with increased zinc
    TITLE OF INVENTION: binding
    NUMBER OF SEQUENCES: 18
    CORRESPONDENCE ADDRESS:
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ADDRESSEE: Finnegan, Henderson, Farabow, Garrett,
      ADDRESSEE: & Dunner, L.L.P.
      STREET: 1300 I Street, N.W.
      CITY: Washington
      STATE: District of Columbia
      COUNTRY: U.S.A.
      ZIP: 20005-3315
    COMPUTER READABLE FORM:
      MEDIUM TYPE: Floppy disk
      COMPUTER: IBM PC compatible
      OPERATING SYSTEM: PC-DOS/MS-DOS
      SOFTWARE: PatentIn Release #1.0, Version #1.25 (EPO)
    CURRENT APPLICATION DATA:
      APPLICATION NUMBER: US/08/900,574
      FILING DATE: July 24, 1997
      CLASSIFICATION: 514
    PRIOR APPLICATION DATA:
      APPLICATION NUMBER: German Application No. 6221837 19630242.0
      FILING DATE: July 26, 1996
    ATTORNEY/AGENT INFORMATION:
      NAME: Carol P. Einaudi
      REGISTRATION NUMBER: 32,220
      REFERENCE/DOCKET NUMBER: 02481.1499-00000
    TELECOMMUNICATION INFORMATION:
      TELEPHONE: (202) 408-4000
      TELEFAX: (202) 408-4400
  INFORMATION FOR SEQ ID NO: 5:
    SEQUENCE CHARACTERISTICS:
      LENGTH: 66 amino acids
      TYPE: Amino acid
      STRANDEDNESS: Single
      TOPOLOGY: linear
    MOLECULE TYPE: Protein
    ORIGINAL SOURCE:
      ORGANISM: Escherichia coli
    FEATURE:
      NAME/KEY: Protein
      LOCATION: 1..66
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Search completed: July 15, 2004, 16:42:32 Job time: 16.1698 secs

### GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 15, 2004, 16:29:19; Search time 11.9776 Seconds

(without alignments)

859.311 Million cell updates/sec

Title: US-09-423-100-6

Perfect score: 587

Sequence: 1 MFPTIPLSRLFDNAMLRAHR.....IVEQCCTSICSLYQLENYCN 107

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Searched: 283366 seqs, 96191526 residues

Total number of hits satisfying chosen parameters: 283366

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

1: pir1:^ 2: pir2:\*

3: pir3:\*

1. pirs.

4: pir4:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

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1	275	46.8	96	2	PC7082	epidermal growth f
2	273.5	46.6	51	1	INWHP	insulin - sperm wh
3	273.5	46.6	51	1	INWHF	insulin - finback
4	273.5	46.6	51	1	INEL	insulin - elephant
5	273	46.5	110	2	B42179	insulin precursor
6	273	46.5	110	2	JQ0178	insulin precursor
7	271.5	46.3	51	1	INHY	insulin - hamster
8	268.5	45.7	51	1	INMSSP	insulin - Egyptian
9	267.5	45.6	51	2	A59151	insulin precursor
10	267	45.5	110	1	IPHU	insulin precursor
11	267	45.5	110	2	A42179	insulin precursor
12	263.5	44.9	51	1	INWH1S	insulin - sei whal
13	263.5	44.9	51	1	INGT	insulin - goat

1.4	262 5	44.0	C 1	1	INCMA	insulin - Arabian
14	263.5	44.9	51			insulin Alabian insulin precursor
15	263	44.8	84	1	IPPG	
16	263	44.8	110	1	INRB	insulin precursor
17	262.5	44.7	51	1	INCT	insulin - cat
18	262	44.6	110	1	IPDG	insulin precursor
19	261.5	44.5	51	1	INMKSQ	insulin - common s
20	260	44.3	110	2	148166	insulin precursor
21	258.5	44.0	105	1	IPBO	insulin precursor
22	257	43.8	108	2	A39883	insulin precursor
23	256.5	43.7	51	2	JQ0362	insulin - North Am
24	255.5	43.5	217	1	STHU	somatotropin 1 pre
25	255.5	43.5	217	2	167410	somatotropin - rhe
26	252.5	43.0	7 <b>7</b>	1	INSH	insulin precursor
27	252	42.9	86	1	IPHO	insulin precursor
28	251.5	42.8	51	1	INCB	insulin - Chinchil
29	250	42.6	108	1	INMS1	insulin 1 precurso
30	249	42.4	110	1	IPRT1	insulin 1 precurso
31	248.5	42.3	51	1	INGS	insulin - goose
32	248	42.2	110	1	IPRT2	insulin 2 precurso
33	248	42.2	110	1	INMS2	insulin 2 precurso
34	246	41.9	52	2	S44469	insulin I1 - North
35	246	41.9	52	2	S44470	insulin I2 - North
36	245	41.7	103	2	I51221	insulin precursor
37	244.5	41.7	51	1	INPQ	insulin - crested
38	244.5	41.7	51	1	INTK	insulin - turkey (
39	244.5	41.7	51	1	INOS	insulin - ostrich
40	244.5	41.7	51	1	A61129	insulin - black-be
41	244.5	41.7	51	2	A60414	insulin - slider t
42	239.5	40.8	107	1	IPCH	insulin precursor
43	238	40.5	52	2	S61361	insulin - Amphiuma
44	235.5	40.1	51	2	S63590	insulin - duckbill
45	233.5	39.8	81	1	IPDK	insulin precursor
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#### ALIGNMENTS

# RESULT 1

PC7082

epidermal growth factor/single chain insulin fusion protein - Bacillus brevis
(fragment)

C; Species: Bacillus brevis

C;Date: 18-Aug-2000 #sequence\_revision 18-Aug-2000 #text\_change 31-Mar-2003

C; Accession: PC7082; PC7083

R; Koh, M.; Hanagata, H.; Ebisu, S.; Morihara, K.; Takagi, H.

Biosci. Biotechnol. Biochem. 64, 1079-1081, 2000

A; Title: Use of Bacillus brevis for synthesis and secretion of Des-B30 single-chain human insulin precursor.

A; Reference number: PC7082; MUID:20335834; PMID:10879487

A;Accession: PC7082 A;Molecule type: DNA A;Residues: 1-96 <KOH> A;Accession: PC7083 A;Molecule type: protein A;Residues: 19-28 <KO2>

C;Genetics: A;Gene: egf-sci

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C; Superfamily: insulin
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Qу
             Db
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RESULT 2
TNWHP
insulin - sperm whale
C; Species: Physeter catodon (sperm whale)
C;Date: 31-Mar-1992 #sequence revision 31-Mar-1992 #text change 16-Jul-1999
C; Accession: A93142; A90082
R; Ishihara, Y.; Saito, T.; Ito, Y.; Fujino, M.
Nature 181, 1468-1469, 1958
A; Title: Structure of sperm- and sei-whale insulins and their breakdown by whale
pepsin.
A; Reference number: A93142
A; Accession: A93142
A; Molecule type: protein
A; Residues: 1-30;31-51 <ISH>
R; Harris, J.I.; Sanger, F.; Naughton, M.A.
Arch. Biochem. Biophys. 65, 427-428, 1956
A; Title: Species differences in insulin.
A; Reference number: A90082
A; Accession: A90082
A; Molecule type: protein
A; Residues: 1-30;31-51 <HAR>
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                        96.2%; Pred. No. 8.7e-22;
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             Db
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RESULT 3
INWHF
insulin - finback whale (tentative sequence)
C; Species: Balaenoptera physalus (finback whale, common rorqual)
C;Date: 31-Mar-1992 #sequence revision 31-Mar-1992 #text change 16-Jul-1999
C; Accession: A91918
R; Hama, H.; Titani, K.; Sakaki, S.; Narita, K.
J. Biochem. 56, 285-293, 1964
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A; Title: The amino acid sequence in fin-whale insulin.

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A; Reference number: A91918
A; Accession: A91918
A; Molecule type: protein
A; Residues: 1-30; 31-51 < HAM>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
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F;31-51/Domain: insulin chain A #status experimental <ACH>
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Qу
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCTSICSLYQLENYCN 51
RESULT 4
INEL
insulin - elephant
C; Species: Elephantidae gen. sp. (elephant)
C; Date: 24-Apr-1984 #sequence revision 30-Sep-1988 #text change 16-Jul-1999
C; Accession: A01584
R; Smith, L.F.
Am. J. Med. 40, 662-666, 1966
A; Title: Species variation in the amino acid sequence of insulin.
A; Reference number: A90029; MUID: 66160119; PMID: 5949593
A; Accession: A01584
A; Molecule type: protein
A; Residues: 1-30;31-51 <SMI>
A; Note: the species of elephant is not given, but it is most probably the Indian
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C; Superfamily: insulin
C; Keywords: hormone; pancreas
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           56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              Db
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RESULT 5
B42179
insulin precursor - green monkey
C; Species: Cercopithecus aethiops (green monkey, grivet)
C;Date: 04-Mar-1993 #sequence revision 18-Nov-1994 #text change 16-Jul-1999
C; Accession: B42179; A05232; S16494; S22056
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R; Seino, S.; Bell, G.I.; Li, W.H.
Mol. Biol. Evol. 9, 193-203, 1992
A; Title: Sequences of primate insulin genes support the hypothesis of a slower
rate of molecular evolution in humans and apes than in monkeys.
A; Reference number: A42179; MUID: 92219953; PMID: 1560757
A; Accession: B42179
A; Molecule type: DNA
A; Residues: 1-110 <SEI>
A; Cross-references: EMBL: X61092; NID: q22808; PIDN: CAA43405.1; PID: q22809
A; Note: sequence extracted from NCBI backbone (NCBIN:95185, NCBIP:95194)
R; Peterson, J.D.; Nehrlich, S.; Oyer, P.E.; Steiner, D.F.
J. Biol. Chem. 247, 4866-4871, 1972
A; Title: Determination of the amino acid sequence of the monkey, sheep, and dog
proinsulin C-peptides by a semi-micro Edman degradation procedure.
A; Reference number: A92111; MUID: 72258016; PMID: 4626369
A; Accession: A05232
A; Molecule type: protein
A; Residues: 57-87 < PET>
C; Genetics:
A; Introns: 63/1
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-24/Domain: signal sequence #status predicted <SIG>
F;25-54/Domain: insulin chain B #status predicted <BCH>
F;25-54,90-110/Product: insulin #status predicted <MAT>
F;57-87/Domain: connecting peptide #status experimental <CPEP>
F;90-110/Domain: insulin chain A #status predicted <ACH>
F;31-96,43-109,95-100/Disulfide bonds: #status predicted
                         46.5%; Score 273; DB 2; Length 110;
  Query Match
                         60.2%; Pred. No. 2.2e-21;
  Best Local Similarity
 Matches 53; Conservative
                               0; Mismatches
                                                1; Indels
                                                               34; Gaps
                                                                            1:
           54 PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
              23 PAFVNQHLCGSHLVEALYLVCGERGFFYTPKTRREAEDPQVGQVELGGGPGAGSLQPLAL 82
Db
           86 ----RGIVEQCCTSICSLYQLENYCN 107
Qу
                    Db
           83 EGSLQKRGIVEQCCTSICSLYQLENYCN 110
RESULT 6
JQ0178
insulin precursor - crab-eating macaque
C; Species: Macaca fascicularis (crab-eating macaque)
C;Date: 07-Sep-1990 #sequence revision 07-Sep-1990 #text change 16-Jul-1999
C; Accession: JQ0178
R; Wetekam, W.; Groneberg, J.; Leineweber, M.; Wengenmayer, F.; Winnacker, E.L.
Gene 19, 179-183, 1982
A; Title: The nucleotide sequence of cDNA coding for preproinsulin from the
primate Macaca fascicularis.
A; Reference number: JQ0178; MUID: 83080474; PMID: 6184262
A; Accession: JQ0178
A; Molecule type: mRNA
A; Residues: 1-110 <WET>
A;Cross-references: GB:J00336; NID:q342121; PIDN:AAA36849.1; PID:q342122
```

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C; Superfamily: insulin
F;1-24/Domain: signal sequence #status predicted <SIG>
F;25-54,90-110/Product: insulin #status predicted <MAT>
F;25-54/Domain: insulin chain B #status predicted <BCH>
F;55-89/Domain: insulin connecting C peptide #status predicted <CPT>
F;90-110/Domain: insulin chain A #status predicted <ACH>
F;31-96,43-109,95-100/Disulfide bonds: #status predicted
                                Score 273; DB 2; Length 110;
 Query Match
                        46.5%;
                        60.2%; Pred. No. 2.2e-21;
 Best Local Similarity
                               0; Mismatches
          53; Conservative
                                                1;
                                                   Indels
                                                             34; Gaps
                                                                         1:
          54 PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
             23 PAFVNOHLCGSHLVEALYLVCGERGFFYTPKTRREAEDPQVGQVELGGGPGAGSLQPLAL 82
Db
          86 -----RGIVEQCCTSICSLYQLENYCN 107
Qу
                   111111111111111111111111
          83 EGSLQKRGIVEQCCTSICSLYQLENYCN 110
Db
RESULT 7
INHY
insulin - hamster
C; Species: Cricetinae gen. sp. (hamster)
C;Date: 31-Mar-1992 #sequence revision 31-Mar-1992 #text_change 16-Jul-1999
C; Accession: A91456
R; Neelon, F.A.; Delcher, H.K.; Steinman, H.; Lebovitz, H.E.
Fed. Proc. 32, 300, 1973
A; Title: Structure of hamster insulin: comparison with a tumor insulin.
A; Reference number: A91456
A; Accession: A91456
A; Molecule type: protein
A; Residues: 1-30;31-51 < NEE>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F:1-30/Domain: insulin chain B #status experimental <BCH>
F;1-30,31-51/Product: insulin #status experimental <MAT>
F;31-51/Domain: insulin chain A #status experimental <ACH>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
 Query Match
                         46.3%; Score 271.5; DB 1;
                                                    Length 51;
 Best Local Similarity
                        94.2%; Pred. No. 1.4e-21;
                                                              1; Gaps
                                                                         1;
 Matches
          49; Conservative
                              2; Mismatches
                                                0; Indels
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKS-GIVDQCCTSICSLYQLENYCN 51
Db
RESULT 8
INMSSP
insulin - Egyptian spiny mouse (tentative sequence)
C; Species: Acomys cahirinus (Egyptian spiny mouse)
C;Date: 13-Jul-1981 #sequence revision 13-Jul-1981 #text_change 31-Mar-2000
C; Accession: A01591
R; Buenzli, H.F.; Humbel, R.E.
```

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Hoppe-Seyler's Z. Physiol. Chem. 353, 444-450, 1972
A; Title: Isolation and partial structural analysis of insulin from mouse (Mus
musculus) and spiny mouse (Acomys cahirinus).
A; Reference number: A01591; MUID: 72189454; PMID: 5028210
A; Contents: composition
A; Accession: A01591
A; Molecule type: protein
A; Residues: 1-30;31-51 <BUE>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-30/Domain: insulin chain B #status predicted <BCH>
F;1-30,31-51/Product: insulin #status predicted <MAT>
F;31-51/Domain: insulin chain A #status predicted <ACH>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
 Query Match
                         45.7%; Score 268.5; DB 1; Length 51;
 Best Local Similarity
                         92.3%; Pred. No. 2.9e-21;
 Matches
          48; Conservative
                                3: Mismatches
                                                 0; Indels
                                                               1; Gaps
                                                                           1;
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             1 FVBQHLCGSHLVEALYLVCGERGFFYTPKS-GIVDQCCTSICSLYQLENYCN 51
Dh
RESULT 9
A59151
insulin precursor - jack bean (fragments)
N; Alternate names: hypoglycemic agent; plant insulin
C; Species: Canavalia ensiformis (jack bean)
C;Date: 07-Dec-1999 #sequence revision 07-Dec-1999 #text change 10-Dec-1999
C; Accession: B59151; A59151
R;Oliveira, A.E.A.; Machado, O.L.T.; Gomes, V.M.; Xavier-Neto, J.; Pereira,
A.C.P.; Vieira, J.G.H.; Fernandes, K.V.S.; Xavier-Filho, J.
Protein Pept. Lett. 6, 15-21, 1999
A; Title: Jack bean seed coat contains a protein with complete sequence homology
to bovine insulin.
A; Reference number: A59151
A; Accession: B59151
A; Molecule type: protein
A; Residues: 1-30 < MACB>
A; Accession: A59151
A; Molecule type: protein
A; Residues: 31-51 < MACA>
C; Comment: The two chains are probably produced from the same precursor.
C; Superfamily: insulin
F;1-30,31-51/Product: insulin #status experimental <MAT>
F;1-30/Domain: chain B #status experimental <CHB>
F;31-51/Domain: chain A #status experimental <CHA>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
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  Query Match
                         92.3%; Pred. No. 3.7e-21;
  Best Local Similarity
  Matches
           48; Conservative
                                1; Mismatches
                                                    Indels
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           56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             Db
           1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASVCSLYQLENYCN 51
```

```
RESULT 10
IPHU
insulin precursor [validated] - human
N; Alternate names: preproinsulin
C; Species: Homo sapiens (man)
C;Date: 23-Oct-1981 #sequence revision 23-Oct-1981 #text change 08-Dec-2000
C; Accession: A93222; A94253; A93216; A94251; A93144; A92075; A91186; I58114;
A01579; S58661
R; Bell, G.I.; Pictet, R.L.; Rutter, W.J.; Cordell, B.; Tischer, E.; Goodman,
H.M.
Nature 284, 26-32, 1980
A; Title: Sequence of the human insulin gene.
A; Reference number: A93222; MUID: 80120725; PMID: 6243748
A; Accession: A93222
A; Molecule type: DNA
A; Residues: 1-110 <BEL>
A;Cross-references: GB:J00265; NID:q186429; PIDN:AAA59172.1; PID:q386828
R; Ullrich, A.; Dull, T.J.; Gray, A.; Brosius, J.; Sures, I.
Science 209, 612-615, 1980
A; Title: Genetic variation in the human insulin gene.
A; Reference number: A94253; MUID: 80236313; PMID: 6248962
A; Accession: A94253
A; Molecule type: DNA
A; Residues: 1-110 <ULL>
A;Cross-references: GB:J00265; NID:g186429; PIDN:AAA59172.1; PID:g386828
R; Bell, G.I.; Swain, W.F.; Pictet, R.; Cordell, B.; Goodman, H.M.; Rutter, W.J.
Nature 282, 525-527, 1979
A; Title: Nucleotide sequence of a cDNA clone encoding human preproinsulin.
A; Reference number: A93216; MUID: 80054779; PMID: 503234
A; Accession: A93216
A; Molecule type: mRNA
A; Residues: 1-110 <BEL2>
A;Cross-references: GB:J00265; NID:g186429; PIDN:AAA59172.1; PID:g386828
R; Sures, I.; Goeddel, D.V.; Gray, A.; Ullrich, A.
Science 208, 57-59, 1980
A; Title: Nucleotide sequence of human preproinsulin complementary DNA.
A; Reference number: A94251; MUID: 80147417; PMID: 6927840
A; Accession: A94251
A; Molecule type: mRNA
A; Residues: 1-110 <SUR>
A;Cross-references: GB:J00265; NID:q186429; PIDN:AAA59172.1; PID:q386828
R; Nicol, D.S.H.W.; Smith, L.F.
Nature 187, 483-485, 1960
A; Title: Amino-acid sequence of human insulin.
A; Reference number: A93144
A; Accession: A93144
A; Molecule type: protein
A; Residues: 25-54; 90-110 < NIC>
R;Oyer, P.E.; Cho, S.; Peterson, J.D.; Steiner, D.F.
J. Biol. Chem. 246, 1375-1386, 1971
A; Title: Studies on human proinsulin. Isolation and amino acid sequence of the
human pancreatic C-peptide.
A; Reference number: A92075; MUID: 71116410; PMID: 5101771
A; Accession: A92075
A; Molecule type: protein
```

```
A; Residues: 57-87 < OYE>
R; Ko, A.; Smyth, D.G.; Markussen, J.; Sundby, F.
Eur. J. Biochem. 20, 190-199, 1971
A; Title: Amino acid sequence of the C-peptide of human proinsulin.
A; Reference number: A91186; MUID: 71257722; PMID: 5560404
A; Accession: A91186
A; Molecule type: protein
A; Residues: 57-87 < KOA>
R; Lucassen, A.M.; Julier, C.; Beressi, J.P.; Boitard, C.; Froquel, P.; Lathrop,
M.; Bell, J.I.
Nature Genet. 4, 305-310, 1993
A; Title: Susceptibility to insulin dependent diabetes mellitus maps to a 4.1 kb
segment of DNA spanning the insulin gene and associated VNTR.
A; Reference number: I58114; MUID: 93364428; PMID: 8358440
A; Accession: I58114
A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-59,63-110 < RES>
A; Cross-references: GB:L15440; NID:q307071; PIDN:AAA59179.1; PID:q307072
R; Sieber, P.; Kamber, B.; Hartmann, A.; Joehl, A.; Riniker, B.; Rittel, W.
Helv. Chim. Acta 57, 2617-2621, 1974
A; Title: Total synthese von Humaninsulin unter gezielter Bildung der
Disulfidbindungen.
A; Reference number: A91636; MUID: 75077277; PMID: 4443293
A; Contents: annotation; synthesis
A; Note: disulfide-bonded human insulin was synthesized; the synthetic hormone
was identical with the natural hormone in chemical and biological activities
A; Note: article in German with English abstract
R; Naithani, V.K.
Hoppe-Seyler's Z. Physiol. Chem. 354, 659-672, 1973
A; Title: The synthesis of C-peptide of human proinsulin.
A; Reference number: A91658; MUID: 75040007; PMID: 4803504
A; Contents: annotation; synthesis of residues 57-87
R; Geiger, R.; Jaeger, G.; Koenig, W.
Chem. Ber. 106, 2347-2352, 1973
A; Title: Synthesis of the complete sequence of human proinsulin C-peptide and
its [Glu-9,Gln-11] analogue.
A; Reference number: A90914
A; Contents: annotation; synthesis of residues 57-87
R; Kaufmann, J.E.; Irminger, J.C.; Halban, P.A.
Biochem. J. 310, 869-874, 1995
A; Title: Sequence requirements for proinsulin processing at the B-chain/C-
peptide junction.
A; Reference number: S58661; MUID: 96013185; PMID: 7575420
A; Contents: annotation; site-directed mutagenesis study of proteolytic
processing
C; Genetics:
A; Gene: GDB: INS
A; Cross-references: GDB:119349; OMIM:176730
A; Map position: 11p15.5-11p15.5
A; Introns: 63/1
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-24/Domain: signal sequence #status predicted <SIG>
F;25-54/Domain: insulin chain B #status experimental <BCH>
F;25-54,90-110/Product: insulin #status experimental <MAT>
F;57-87/Domain: connecting C peptide #status experimental <CPEP>
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F;31-96,43-109,95-100/Disulfide bonds: #status experimental
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 Query Match
 Best Local Similarity 60.5%; Pred. No. 9.3e-21;
 Matches
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                                                                     1:
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Qv
            Db
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Qy
          86 ----RGIVEQCCTSICSLYQLENYCN 107
                Db
          85 SLQKRGIVEQCCTSICSLYQLENYCN 110
RESULT 11
A42179
insulin precursor - chimpanzee
C; Species: Pan troglodytes (chimpanzee)
C;Date: 04-Mar-1993 #sequence revision 18-Nov-1994 #text change 16-Jul-1999
C; Accession: A42179; S22058
R; Seino, S.; Bell, G.I.; Li, W.H.
Mol. Biol. Evol. 9, 193-203, 1992
A; Title: Sequences of primate insulin genes support the hypothesis of a slower
rate of molecular evolution in humans and apes than in monkeys.
A; Reference number: A42179; MUID: 92219953; PMID: 1560757
A; Accession: A42179
A; Status: preliminary
A; Molecule type: DNA
A; Residues: 1-110 <SEI>
A; Cross-references: EMBL: X61089; NID: q38251; PIDN: CAA43403.1; PID: q38252
A; Note: sequence extracted from NCBI backbone (NCBIP:95067)
C:Genetics:
A; Introns: 63/1
C; Superfamily: insulin
 Query Match
                       45.5%; Score 267; DB 2; Length 110;
 Best Local Similarity 60.5%; Pred. No. 9.3e-21;
 Matches
         52; Conservative 0; Mismatches 0; Indels 34; Gaps
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
             Db
          25 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRREAEDLQVGQVELGGGPGAGSLQPLALEG 84
          86 ----RGIVEQCCTSICSLYQLENYCN 107
Qу
                85 SLQKRGIVEQCCTSICSLYQLENYCN 110
Dh
RESULT 12
INWH1S
insulin - sei whale
C; Species: Balaenoptera borealis (sei whale)
C;Date: 13-Jul-1981 #sequence_revision 13-Jul-1981 #text_change 16-Jul-1999
C; Accession: A01582
R; Ishihara, Y.; Saito, T.; Ito, Y.; Fujino, M.
```

F;90-110/Domain: insulin chain A #status experimental <ACH>

```
Nature 181, 1468-1469, 1958
A; Title: Structure of sperm- and sei-whale insulins and their breakdown by whale
pepsin.
A; Reference number: A93142
A; Accession: A01582
A; Molecule type: protein
A; Residues: 1-30; 31-51 <ISH>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-30/Domain: insulin chain B #status experimental <BCH>
F;1-30,31-51/Product: insulin #status experimental <MAT>
F;31-51/Domain: insulin chain A #status experimental <ACH>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
 Query Match
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 Best Local Similarity
 Matches
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                                                 3; Indels
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Qу
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
             Db
           1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASTCSLYQLENYCN 51
RESULT 13
INGT
insulin - goat
C; Species: Capra aegagrus hircus (domestic goat)
C; Date: 13-Jul-1981 #sequence revision 13-Jul-1981 #text change 16-Jul-1999
C; Accession: A01586
R; Smith, L.F.
Am. J. Med. 40, 662-666, 1966
A; Title: Species variation in the amino acid sequence of insulin.
A; Reference number: A90029; MUID: 66160119; PMID: 5949593
A; Accession: A01586
A; Molecule type: protein
A; Residues: 1-30; 31-51 <SMI>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-30/Domain: insulin chain B #status experimental <BCH>
F;1-30,31-51/Product: insulin #status experimental <MAT>
F;31-51/Domain: insulin chain A #status experimental <ACH>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
 Query Match
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 Best Local Similarity
                        90.4%; Pred. No. 9.6e-21;
 Matches
           47; Conservative
                               1; Mismatches
                                                 3; Indels
                                                              1; Gaps
                                                                          1:
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             Db
           1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCAGVCSLYQLENYCN 51
RESULT 14
INCMA
insulin - Arabian camel (tentative sequence)
C; Species: Camelus dromedarius (Arabian camel)
C;Date: 31-Mar-1992 #sequence revision 31-Mar-1992 #text_change 16-Jul-1999
```

```
C; Accession: A92782
R; Danho, W.O.
J. Fac. Med. Baghdad 14, 16-28, 1972
A; Title: The isolation and characterization of insulin of camel (Camelus
dromedarius).
A; Reference number: A92782
A; Accession: A92782
A; Molecule type: protein
A; Residues: 1-30; 31-51 < DAN>
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-30/Domain: insulin chain B #status experimental <BCH>
F;1-30,31-51/Product: insulin #status experimental <MAT>
F;31-51/Domain: insulin chain A #status experimental <ACH>
F;7-37,19-50,36-41/Disulfide bonds: #status predicted
  Query Match
                          44.9%; Score 263.5; DB 1; Length 51;
                          90.4%; Pred. No. 9.6e-21;
  Best Local Similarity
  Matches
            47; Conservative
                                 1; Mismatches
                                                   3; Indels
                                                                 1; Gaps
                                                                              1;
Qу
           56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
              Db
            1 FANQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASVCSLYQLENYCN 51
RESULT 15
IPPG
insulin precursor - piq
C; Species: Sus scrofa domestica (domestic pig)
C;Date: 22-Jun-1981 #sequence revision 22-Jun-1981 #text change 16-Jul-1999
C; Accession: A01583; A94572; S16492; A60835; B60835
R; Chance, R.E.; Ellis, R.M.; Bromer, W.W.
Science 161, 165-167, 1968
A; Title: Porcine proinsulin: characterization and amino acid sequence.
A; Reference number: A94240; MUID: 68286485; PMID: 5657063
A; Accession: A01583
A; Molecule type: protein .
A; Residues: 1-34, 'Q', 36-84 < CHA>
R; Chance, R.E.
submitted to the Atlas, July 1970
A; Reference number: A94572
A; Accession: A94572
A; Molecule type: protein
A; Residues: 1-84 <CH2>
R;Brown, H.; Sanger, F.; Kitai, R.
Biochem. J. 60, 556-565, 1955
A; Title: The structure of pig and sheep insulins.
A; Reference number: A90344
A; Accession: S16492
A; Molecule type: protein
A; Residues: 1-30; 31-51 < BRO>
R; Snel, L.; Damgaard, U.
Horm. Metab. Res. 20, 476-480, 1988
A; Title: Proinsulin heterogeneity in pigs.
A; Reference number: A60835; MUID: 89032178; PMID: 3181865
A; Accession: A60835
A; Molecule type: protein
```

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A; Residues: 33-38,40-62 <SNE>
A; Note: the authors report the characterization of a connecting peptide variant
lacking Ala-39
A; Accession: B60835
A; Molecule type: protein
A; Residues: 33-62 <SN2>
R; Blundell, T.; Dodson, G.; Hodgkin, D.; Mercola, D.
Adv. Protein Chem. 26, 279-402, 1972
A; Title: Insulin. the structure in the crystal and its reflection in chemistry
and biology.
A; Reference number: A90017
A; Contents: annotation; X-ray crystallography, 1.9 angstroms
C; Superfamily: insulin
C; Keywords: hormone; pancreas
F;1-30/Domain: insulin chain B #status experimental <BCH>
F;1-30,64-84/Product: insulin #status experimental <MAT>
F;33-63/Domain: connecting peptide #status experimental <CPEP>
F;64-84/Domain: insulin chain A #status experimental <ACH>
F;7-70,19-83,69-74/Disulfide bonds: #status experimental
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Qу
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Db
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Qу
               Db
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Search completed: July 15, 2004, 16:37:33

Job time : 12.1443 secs

### GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 15, 2004, 16:37:41; Search time 44.5168 Seconds

(without alignments)

751.267 Million cell updates/sec

Title: US-09-423-100-6

Perfect score: 587

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Scoring table: BLOSUM62

Gapop 10.0, Gapext 0.5

Searched: 1285345 seqs, 312560633 residues

Total number of hits satisfying chosen parameters: 1285345

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: Published Applications AA:\*

1: /cgn2\_6/ptodata/1/pubpaa/US07\_PUBCOMB.pep:\*

2: /cgn2\_6/ptodata/1/pubpaa/PCT\_NEW\_PUB.pep:\*

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5: /cgn2 6/ptodata/1/pubpaa/US07 NEW PUB.pep:\*

6: /cgn2 6/ptodata/1/pubpaa/PCTUS\_PUBCOMB.pep:\*

: /cgn2 6/ptodata/1/pubpaa/US08 NEW PUB.pep:\*

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16: /cgn2\_6/ptodata/1/pubpaa/US10 NEW PUB.pep:\*

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Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

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Result Query

No. Score Match Length DB ID

Description

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2	555.5	94.6	150	13	US-10-054-873-7	Sequence 7, Appli
3	302.5	51.5	137	16	US-10-101-454-39	Sequence 39, Appl
4	299	50.9	145	16	US-10-101-454-45	Sequence 45, Appl
5	299	50.9	146	16	US-10-101-454-48	Sequence 48, Appl
6	294	50.1	52	13	US-10-054-873-5	Sequence 5, Appli
7	284.5	48.5	124	9	US-09-894-711-18	Sequence 18, Appl
8	284	48.4	138	9	US-09-861-687-19	Sequence 19, Appl
9	284	48.4	138	12	US-10-620-651-19	Sequence 19, Appl
10	284	48.4	140	16	US-10-101-454-33	Sequence 33, Appl
11	284	48.4	140	16	US-10-101-454-42	Sequence 42, Appl
12	281	47.9	104	16	US-10-101-454-15	Sequence 15, Appl
13	278.5	47.4	51	10	US-09-858-935B-5	Sequence 5, Appli
14	278.5	47.4	51	12	US-10-444-649-3	Sequence 3, Appli
15	278.5	47.4	51	12	US-10-444-701-3	Sequence 3, Appli
16	278.5	47.4	51	12	US-10-271-869-5	Sequence 5, Appli
17	278.5	47.4	51	13	US-10-028-410-3	Sequence 3, Appli
18	278.5	47.4	51	14	US-10-444-326-3	Sequence 3, Appli-
19	278.5	47.4	51	16	US-10-444-262-3	Sequence 3, Appli
20	278	47.4	117	9	US-09-280-030-63	Sequence 63, Appl
21	277.5	47.3	124	15	US-10-221-677-24	Sequence 24, Appl
22	277	47.2	96	9	US-09-947-563-4	Sequence 4, Appli
23	277	47.2	102	16	US-10-101-454-36	Sequence 36, Appl
24	275.5	46.9	124	9	US-09-736-611-12	Sequence 12, Appl
25	275.5	46.9	124	9	US-09-740-359-12	Sequence 12, Appl
26	275.5	46.9	124	9	US-09-894-711-12	Sequence 12, Appl
27	275.5	46.9	124	14	US-10-316-421-12	Sequence 12, Appl
28	275.5	46.9	125	9	US-09-736-611-10	Sequence 10, Appl
29	275.5	46.9	125	9	US-09-740-359-10	Sequence 10, Appl
30	275.5	46.9	125	9	US-09-894-711-10	Sequence 10, Appl
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32	275.5	46.9	147	9	US-09-736-611-8	Sequence 8, Appli
33	275.5	46.9	147	9	US-09-740-359-7	Sequence 7, Appli
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35	274	46.7	144	9	US-09-736-611-6	Sequence 6, Appli
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41	270	46.0	104	16	US-10-101-454-21	Sequence 21, Appl
42	270	46.0	104	16	US-10-101-454-27	Sequence 27, Appl
43	269.5	45.9	130	9	US-09-280-030-62	Sequence 62, Appl
44	269	45.8	104	16	US-10-101-454-24	Sequence 24, Appl
45	269	45.8	104	16	US-10-101-454-30	Sequence 30, Appl

## ALIGNMENTS

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RESULT 1
US-10-054-873-6
; Sequence 6, Application US/10054873
; Publication No. US20020164712A1
; GENERAL INFORMATION:
; APPLICANT: Gan, Zhong Ru
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TITLE OF INVENTION: Chimeric Protein Containing an
;
                           Intramolecular Chaperone-Like Sequence
        NUMBER OF SEQUENCES: 7
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: Townsend and Townsend and Crew LLP
             STREET: Two Embarcadero Center, Eighth Floor
             CITY: San Francisco
             STATE: California
             COUNTRY: USA
             ZIP: 94111-3834
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.30
        CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/10/054,873
             FILING DATE: 22-Jan-2002
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
             APPLICATION NUMBER: WO PCT/CN98/00052
             FILING DATE: 31-MAR-1998
             APPLICATION NUMBER: US 09/423,100
             FILING DATE: 11-DEC-2000
        ATTORNEY/AGENT INFORMATION:
             NAME: Mycroft, Frank J
             REGISTRATION NUMBER: 46,946
             REFERENCE/DOCKET NUMBER: 020167-000130US
   INFORMATION FOR SEQ ID NO: 6:
        SEQUENCE CHARACTERISTICS:
             LENGTH: 107 amino acids
             TYPE: amino acid
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             TOPOLOGY: linear
        MOLECULE TYPE: protein
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US-10-054-873-7
; Sequence 7, Application US/10054873
; Publication No. US20020164712A1
    GENERAL INFORMATION:
        APPLICANT: Gan, Zhong Ru
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TITLE OF INVENTION: Chimeric Protein Containing an
                          Intramolecular Chaperone-Like Sequence
       NUMBER OF SEQUENCES: 7
       CORRESPONDENCE ADDRESS:
            ADDRESSEE: Townsend and Townsend and Crew LLP
            STREET: Two Embarcadero Center, Eighth Floor
            CITY: San Francisco
            STATE: California
            COUNTRY: USA
            ZIP: 94111-3834
       COMPUTER READABLE FORM:
            MEDIUM TYPE: Floppy disk
            COMPUTER: IBM PC compatible
            OPERATING SYSTEM: PC-DOS/MS-DOS
            SOFTWARE: PatentIn Release #1.0, Version #1.30
       CURRENT APPLICATION DATA:
            APPLICATION NUMBER: US/10/054,873
            FILING DATE: 22-Jan-2002
            CLASSIFICATION: <Unknown>
       PRIOR APPLICATION DATA:
            APPLICATION NUMBER: WO PCT/CN98/00052
            FILING DATE: 31-MAR-1998
            APPLICATION NUMBER: US 09/423,100
            FILING DATE: 11-DEC-2000
       ATTORNEY/AGENT INFORMATION:
            NAME: Mycroft, Frank J
            REGISTRATION NUMBER: 46,946
            REFERENCE/DOCKET NUMBER: 020167-000130US
 INFORMATION FOR SEQ ID NO: 7:
       SEQUENCE CHARACTERISTICS:
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            STRANDEDNESS: <Unknown>
            TOPOLOGY: linear
       MOLECULE TYPE: protein
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US-10-054-873-7
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; Sequence 39, Application US/10101454
; Publication No. US20040110664A1
   GENERAL INFORMATION:
        APPLICANT: Havelund, Svend
                   Halstrom, John
                   Jonassen, Ib
                   Andersen, Asser Sloth
                   Markussen, Jan
        TITLE OF INVENTION: ACYLATED INSULIN
        NUMBER OF SEQUENCES: 49
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: Novo Nordisk of North America, Inc.
             STREET: 405 Lexington Avenue, 64th Floor
             CITY: New York
             STATE: New York
             COUNTRY: United States of America
             ZIP: 10174-6401
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.25
        CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/10/101,454
             FILING DATE: 20-Mar-2002
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
             APPLICATION NUMBER: 08/400,256
             FILING DATE: 03-MAR-1995
        ATTORNEY/AGENT INFORMATION:
             NAME: Lambiris, Elias J.
             REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 3985.220-US
        TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
   INFORMATION FOR SEQ ID NO: 39:
        SEQUENCE CHARACTERISTICS:
             LENGTH: 137 amino acids
             TYPE: amino acid
             TOPOLOGY: linear
        MOLECULE TYPE: protein
        SEQUENCE DESCRIPTION: SEQ ID NO: 39:
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Db
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; Sequence 45, Application US/10101454
; Publication No. US20040110664A1
    GENERAL INFORMATION:
         APPLICANT: Havelund, Svend
                    Halstrom, John
;
                    Jonassen, Ib
                    Andersen, Asser Sloth
                    Markussen, Jan
         TITLE OF INVENTION: ACYLATED INSULIN
         NUMBER OF SEQUENCES: 49
         CORRESPONDENCE ADDRESS:
              ADDRESSEE: Novo Nordisk of North America, Inc.
              STREET: 405 Lexington Avenue, 64th Floor
              CITY: New York
              STATE: New York
              COUNTRY: United States of America
              ZIP: 10174-6401
         COMPUTER READABLE FORM:
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              COMPUTER: IBM PC compatible
              OPERATING SYSTEM: PC-DOS/MS-DOS
              SOFTWARE: PatentIn Release #1.0, Version #1.25
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              FILING DATE: 20-Mar-2002
              CLASSIFICATION: <Unknown>
         PRIOR APPLICATION DATA:
              APPLICATION NUMBER: 08/400,256
              FILING DATE: 03-MAR-1995
         ATTORNEY/AGENT INFORMATION:
              NAME: Lambiris, Elias J.
              REGISTRATION NUMBER: 33,728
              REFERENCE/DOCKET NUMBER: 3985.220-US
         TELECOMMUNICATION INFORMATION:
              TELEPHONE: 212-867-0123
              TELEFAX: 212-878-9655
    INFORMATION FOR SEQ ID NO: 45:
         SEQUENCE CHARACTERISTICS:
              LENGTH: 145 amino acids
              TYPE: amino acid
              TOPOLOGY: linear
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US-10-101-454-48
; Sequence 48, Application US/10101454
; Publication No. US20040110664A1
    GENERAL INFORMATION:
         APPLICANT: Havelund, Svend
                    Halstrom, John
                    Jonassen, Ib
                    Andersen, Asser Sloth
                    Markussen, Jan
         TITLE OF INVENTION: ACYLATED INSULIN
         NUMBER OF SEQUENCES: 49
         CORRESPONDENCE ADDRESS:
              ADDRESSEE: Novo Nordisk of North America, Inc.
              STREET: 405 Lexington Avenue, 64th Floor
              CITY: New York
              STATE: New York
              COUNTRY: United States of America
              ZIP: 10174-6401
         COMPUTER READABLE FORM:
              MEDIUM TYPE: Floppy disk
              COMPUTER: IBM PC compatible
              OPERATING SYSTEM: PC-DOS/MS-DOS
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              FILING DATE: 20-Mar-2002
              CLASSIFICATION: <Unknown>
         PRIOR APPLICATION DATA:
              APPLICATION NUMBER: 08/400,256
              FILING DATE: 03-MAR-1995
         ATTORNEY/AGENT INFORMATION:
              NAME: Lambiris, Elias J.
              REGISTRATION NUMBER: 33,728
              REFERENCE/DOCKET NUMBER: 3985.220-US
         TELECOMMUNICATION INFORMATION:
              TELEPHONE: 212-867-0123
              TELEFAX: 212-878-9655
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         SEQUENCE CHARACTERISTICS:
              LENGTH: 146 amino acids
              TYPE: amino acid
              TOPOLOGY: linear
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; Publication No. US20020164712A1
   GENERAL INFORMATION:
        APPLICANT: Gan, Zhong Ru
        TITLE OF INVENTION: Chimeric Protein Containing an
                            Intramolecular Chaperone-Like Sequence
        NUMBER OF SEQUENCES: 7
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: Townsend and Townsend and Crew LLP
             STREET: Two Embarcadero Center, Eighth Floor
             CITY: San Francisco
             STATE: California
             COUNTRY: USA
             ZIP: 94111-3834
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.30
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             FILING DATE: 22-Jan-2002
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
             APPLICATION NUMBER: WO PCT/CN98/00052
             FILING DATE: 31-MAR-1998
             APPLICATION NUMBER: US 09/423,100
             FILING DATE: 11-DEC-2000
        ATTORNEY/AGENT INFORMATION:
             NAME: Mycroft, Frank J
             REGISTRATION NUMBER: 46,946
             REFERENCE/DOCKET NUMBER: 020167-000130US
   INFORMATION FOR SEQ ID NO: 5:
        SEQUENCE CHARACTERISTICS:
             LENGTH: 52 amino acids
             TYPE: amino acid
             STRANDEDNESS: <Unknown>
             TOPOLOGY: linear
        MOLECULE TYPE: protein
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; Sequence 18, Application US/09894711
; Patent No. US20020137144A1
; GENERAL INFORMATION:
; APPLICANT: Kjeldsen, Thomas Borglum
  APPLICANT: Ludvigsen, Svend
  TITLE OF INVENTION: Method for making insulin precursors and
  TITLE OF INVENTION: insulin precursor analogues having improved fermentation
  TITLE OF INVENTION: yield in yeast
  FILE REFERENCE: 6148.400-US
  CURRENT APPLICATION NUMBER: US/09/894,711
  CURRENT FILING DATE: 2001-06-28
  PRIOR APPLICATION NUMBER: PA 2000 00443
  PRIOR FILING DATE: 2000-03-17
  PRIOR APPLICATION NUMBER: PA 1999 01869
  PRIOR FILING DATE: 1999-12-29
  PRIOR APPLICATION NUMBER: 60/211,081
  PRIOR FILING DATE: 2000-06-13
  PRIOR APPLICATION NUMBER: 60/181,450
; PRIOR FILING DATE: 2000-02-10
; PRIOR APPLICATION NUMBER: 09/740,359
  PRIOR FILING DATE: 2000-12-19
  NUMBER OF SEQ ID NOS: 20
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RESULT 8
US-09-861-687-19
; Sequence 19, Application US/09861687
; Publication No. US20020193292A1
    GENERAL INFORMATION:
         APPLICANT: Markussen, Jan
                   Jonassen, Ib
                   Havelund, Svend
                   Brandt, Jakob
                   Kurtzhals, Peter
                   Hansen, Hertz Per
                   Kaarsholm, Niels Christian
         TITLE OF INVENTION: INSULIN DERIVATIVES
         NUMBER OF SEQUENCES: 26
         CORRESPONDENCE ADDRESS:
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ADDRESSEE: No. US20020193292Alo No. US20020193292Aldisk of No.
US20020193292A1th America, Inc.
            STREET: 405 Lexington Avenue, 64th Floor
             CITY: New York
             STATE: New York
             COUNTRY: United States of America
             ZIP: 10174-6401
        COMPUTER READABLE FORM:
            MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.30
        CURRENT APPLICATION DATA:
            APPLICATION NUMBER: US/09/861,687
             FILING DATE: 21-May-2001
            CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
            APPLICATION NUMBER: 08/932,082
             FILING DATE: 16-DEC-1997
        ATTORNEY/AGENT INFORMATION:
            NAME: Lambiris, Elias J.
             REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 4341.204-US
        TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
   INFORMATION FOR SEQ ID NO: 19:
        SEQUENCE CHARACTERISTICS:
             LENGTH: 138 amino acids
             TYPE: amino acid
             TOPOLOGY: linear
        MOLECULE TYPE: protein
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US-09-861-687-19
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  Best Local Similarity 48.2%; Pred. No. 9.8e-26;
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Qу
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; Sequence 19, Application US/10620651
; Publication No. US20040067874A1
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APPLICANT: Markussen, Jan
                  Jonassen, Ib
                  Havelund, Svend
                  Brandt, Jakob
                  Kurtzhals, Peter
                  Hansen, Hertz Per
                  Kaarsholm, Niels Christian
        TITLE OF INVENTION: INSULIN DERIVATIVES
        NUMBER OF SEQUENCES: 26
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: No. US20040067874Alo No. US20040067874Aldisk of No.
US20040067874A1th America, Inc.
             STREET: 405 Lexington Avenue, 64th Floor
;
             CITY: New York
;
             STATE: New York
             COUNTRY: United States of America
             ZIP: 10174-6401
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.30
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             FILING DATE: 16-Jul-2003
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
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             FILING DATE: 17-SEPT-1997
        ATTORNEY/AGENT INFORMATION:
             NAME: Lambiris, Elias J.
             REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 4341.204-US
        TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
   INFORMATION FOR SEQ ID NO: 19:
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             TOPOLOGY: linear
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          118 GIVEQCCTSICSLYQLENYCN 138
RESULT 10
US-10-101-454-33
; Sequence 33, Application US/10101454
; Publication No. US20040110664A1
    GENERAL INFORMATION:
        APPLICANT: Havelund, Svend
                   Halstrom, John
                   Jonassen, Ib
                   Andersen, Asser Sloth
                   Markussen, Jan
         TITLE OF INVENTION: ACYLATED INSULIN
        NUMBER OF SEQUENCES: 49
         CORRESPONDENCE ADDRESS:
             ADDRESSEE: Novo Nordisk of North America, Inc.
              STREET: 405 Lexington Avenue, 64th Floor
             CITY: New York
              STATE: New York
             COUNTRY: United States of America
              ZIP: 10174-6401
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
              COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
              SOFTWARE: PatentIn Release #1.0, Version #1.25
         CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/10/101,454
              FILING DATE: 20-Mar-2002
             CLASSIFICATION: <Unknown>
         PRIOR APPLICATION DATA:
             APPLICATION NUMBER: 08/400,256
              FILING DATE: 03-MAR-1995
        ATTORNEY/AGENT INFORMATION:
             NAME: Lambiris, Elias J.
              REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 3985.220-US
         TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
    INFORMATION FOR SEQ ID NO: 33:
         SEQUENCE CHARACTERISTICS:
              LENGTH: 140 amino acids
              TYPE: amino acid
             TOPOLOGY: linear
        MOLECULE TYPE: protein
         SEQUENCE DESCRIPTION: SEQ ID NO: 33:
US-10-101-454-33
  Query Match
                         48.4%; Score 284; DB 16; Length 140;
                         47.6%; Pred. No. 9.9e-26;
  Best Local Similarity
                                6; Mismatches 27; Indels
  Matches
           68; Conservative
                                                               42; Gaps
                                                                            5;
           2 FPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQ--KYSFLQ-----N 48
Qу
```

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11:1 : 1
                                    3 FPSI----FTAVLFAASSALAAPVNTTTEDETAQIPAEAVIGYSDLEGDFDVAVLPFSN 57
Db
          49 PLGTG-----PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT-- 85
Qу
                Db
          58 STNNGLLFINTTIASIAAKEEGVSLDKRFVNQHLCGSHLVEALYLVCGERGFFYTPKSDD 117
          86 -RGIVEQCCTSICSLYQLENYCN 107
Qy
              Db
         118 AKGIVEQCCTSICSLYQLENYCN 140
RESULT 11
US-10-101-454-42
; Sequence 42, Application US/10101454
; Publication No. US20040110664A1
   GENERAL INFORMATION:
        APPLICANT: Havelund, Svend
                  Halstrom, John
                  Jonassen, Ib
                  Andersen, Asser Sloth
                  Markussen, Jan
        TITLE OF INVENTION: ACYLATED INSULIN
        NUMBER OF SEQUENCES: 49
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: Novo Nordisk of North America, Inc.
             STREET: 405 Lexington Avenue, 64th Floor
             CITY: New York
             STATE: New York
             COUNTRY: United States of America
             ZIP: 10174-6401
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.25
        CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/10/101,454
             FILING DATE: 20-Mar-2002
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
             APPLICATION NUMBER: 08/400,256
             FILING DATE: 03-MAR-1995
        ATTORNEY/AGENT INFORMATION:
             NAME: Lambiris, Elias J.
             REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 3985.220-US
        TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
   INFORMATION FOR SEQ ID NO: 42:
        SEQUENCE CHARACTERISTICS:
             LENGTH: 140 amino acids
             TYPE: amino acid
             TOPOLOGY: linear
        MOLECULE TYPE: protein
        SEQUENCE DESCRIPTION: SEQ ID NO: 42:
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48.4%; Score 284; DB 16; Length 140;
 Query Match
 Best Local Similarity 47.6%; Pred. No. 9.9e-26;
                              6; Mismatches 27; Indels 42; Gaps
 Matches 68; Conservative
                                                                        5;
           2 FPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQ--KYSFLQ-----N 48
Qγ
                   | : |
                                   3 FPSI----FTAVLFAASSALAAPVNTTTEDETAQIPAEAVIGYSDLEGDFDVAVLPFSN 57
          49 PLGTG-----PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT-- 85
Qу
                -1
                                      58 STNNGLLFINTTIASIAAKEEGVSMAKRFVNOHLCGSHLVEALYLVCGERGFFYTPKSDD 117
Qу
          86 -RGIVEQCCTSICSLYQLENYCN 107
             :||||||||
         118 AKGIVEQCCTSICSLYQLENYCN 140
RESULT 12
US-10-101-454-15
; Sequence 15, Application US/10101454
; Publication No. US20040110664A1
   GENERAL INFORMATION:
        APPLICANT: Havelund, Svend
                  Halstrom, John
                  Jonassen, Ib
                  Andersen, Asser Sloth
                  Markussen, Jan
        TITLE OF INVENTION: ACYLATED INSULIN
        NUMBER OF SEQUENCES: 49
        CORRESPONDENCE ADDRESS:
             ADDRESSEE: Novo Nordisk of North America, Inc.
             STREET: 405 Lexington Avenue, 64th Floor
             CITY: New York
             STATE: New York
             COUNTRY: United States of America
             ZIP: 10174-6401
        COMPUTER READABLE FORM:
             MEDIUM TYPE: Floppy disk
             COMPUTER: IBM PC compatible
             OPERATING SYSTEM: PC-DOS/MS-DOS
             SOFTWARE: PatentIn Release #1.0, Version #1.25
        CURRENT APPLICATION DATA:
             APPLICATION NUMBER: US/10/101,454
             FILING DATE: 20-Mar-2002
             CLASSIFICATION: <Unknown>
        PRIOR APPLICATION DATA:
             APPLICATION NUMBER: 08/400,256
             FILING DATE: 03-MAR-1995
        ATTORNEY/AGENT INFORMATION:
             NAME: Lambiris, Elias J.
             REGISTRATION NUMBER: 33,728
             REFERENCE/DOCKET NUMBER: 3985.220-US
        TELECOMMUNICATION INFORMATION:
             TELEPHONE: 212-867-0123
             TELEFAX: 212-878-9655
```

```
INFORMATION FOR SEQ ID NO: 15:
        SEQUENCE CHARACTERISTICS:
;
             LENGTH: 104 amino acids
            TYPE: amino acid
;
            TOPOLOGY: linear
        MOLECULE TYPE: protein
        SEQUENCE DESCRIPTION: SEQ ID NO: 15:
US-10-101-454-15
                        47.9%; Score 281; DB 16; Length 104;
 Query Match
 Best Local Similarity 71.8%; Pred. No. 1.6e-25;
         56; Conservative 6; Mismatches 8; Indels
 Matches
                                                           8; Gaps
                                                                      3:
          37 IPKEQ----KYSFLQNPLGTGPRFVNQHLCGSHLVEALYLVCGERGFFYTPKT---RGIV 89
Qу
                   28 IPEESLIIAENTTLAN-VAMAKRFVNQHLCGSHLVEALYLVCGERGFFYTPKSDDAKGIV 86
          90 EQCCTSICSLYQLENYCN 107
Qу
             111111111111111111
Db
          87 EQCCTSICSLYQLENYCN 104
RESULT 13
US-09-858-935B-5
; Sequence 5, Application US/09858935B
; Publication No. US20030069177A1
; GENERAL INFORMATION:
 APPLICANT: Dubaquie, Yves
; APPLICANT: Filvaroff, Ellen
; APPLICANT: Lowman, Henry B.
; TITLE OF INVENTION: METHOD FOR TREATING CARTILAGE DISORDERS
; FILE REFERENCE: P1794R1
; CURRENT APPLICATION NUMBER: US/09/858,935B
; CURRENT FILING DATE: 2002-07-02
; PRIOR APPLICATION NUMBER: US 60/248,985
  PRIOR FILING DATE: 2000-11-15
  PRIOR APPLICATION NUMBER: US 60/204,490
; PRIOR FILING DATE: 2000-05-16
; NUMBER OF SEQ ID NOS: 153
; SEQ ID NO 5
   LENGTH: 51
   TYPE: PRT
   ORGANISM: Homo sapiens
US-09-858-935B-5
 Query Match
                        47.4%; Score 278.5; DB 10; Length 51;
  Best Local Similarity 98.1%; Pred. No. 1.3e-25;
                              0; Mismatches 0; Indels
                                                           1; Gaps
 Matches
         51; Conservative
                                                                       1;
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKT-GIVEQCCTSICSLYQLENYCN 51
Db
RESULT 14
US-10-444-649-3
; Sequence 3, Application US/10444649
```

```
; Publication No. US20040033951A1
; GENERAL INFORMATION:
  APPLICANT: Dubaquie, Yves
  APPLICANT: Lowman, Henry
  TITLE OF INVENTION: PROTEIN VARIANTS
  FILE REFERENCE: P1712R1
  CURRENT APPLICATION NUMBER: US/10/444,649
  CURRENT FILING DATE: 2003-05-22
  PRIOR APPLICATION NUMBER: US/09/724,479
 PRIOR FILING DATE: 2000-11-28
  PRIOR APPLICATION NUMBER: US/09/477,923
  PRIOR FILING DATE: 2000-01-05
  NUMBER OF SEO ID NOS: 6
; SEQ ID NO 3
   LENGTH: 51
   TYPE: PRT
   ORGANISM: Homo sapiens
US-10-444-649-3
 Query Match
                        47.4%; Score 278.5; DB 12; Length 51;
 Best Local Similarity
                        98.1%; Pred. No. 1.3e-25;
          51; Conservative
                              0; Mismatches
                                               0; Indels
                                                            1; Gaps
                                                                        1;
Qу
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKT-GIVEQCCTSICSLYQLENYCN 51
Db
RESULT 15
US-10-444-701-3
; Sequence 3, Application US/10444701
; Publication No. US20040033952A1
; GENERAL INFORMATION:
  APPLICANT: Dubaquie, Yves
  APPLICANT: Lowman, Henry
  TITLE OF INVENTION: PROTEIN VARIANTS
  FILE REFERENCE: P1712R1
  CURRENT APPLICATION NUMBER: US/10/444,701
; CURRENT FILING DATE: 2003-05-22
; PRIOR APPLICATION NUMBER: US/09/723,866
; PRIOR FILING DATE: 2000-11-28
; PRIOR APPLICATION NUMBER: US/09/477,923
  PRIOR FILING DATE: 2000-01-05
 NUMBER OF SEQ ID NOS: 6
; SEQ ID NO 3
   LENGTH: 51
   TYPE: PRT
   ORGANISM: Homo sapiens
US-10-444-701-3
                               Score 278.5; DB 12; Length 51;
 Query Match
                        47.4%;
                        98.1%; Pred. No. 1.3e-25;
 Best Local Similarity
 Matches
         51; Conservative
                              0; Mismatches
                                              0; Indels
                                                            1; Gaps
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
             Db
           1 FVNQHLCGSHLVEALYLVCGERGFFYTPKT-GIVEQCCTSICSLYQLENYCN 51
```

Search completed: July 15, 2004, 17:05:09 Job time: 45.5168 secs

### GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 15, 2004, 16:29:50; Search time 36.5317 Seconds

(without alignments)

924.141 Million cell updates/sec

Title: US-09-423-100-6

Perfect score: 587

Sequence: 1 MFPTIPLSRLFDNAMLRAHR.....IVEQCCTSICSLYQLENYCN 107

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 1017041 seqs, 315518202 residues

Total number of hits satisfying chosen parameters: 1017041

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: SPTREMBL 25:\*

1: sp\_archea:\*
2: sp\_bacteria:\*
3: sp fungi:\*

4: sp human:\*

5: sp invertebrate:\*

6: sp\_mammal:\*
7: sp mhc:\*

8: sp\_nuic:\*

b. bp\_organerie.

9: sp\_phage:\*

10: sp\_plant:\*

11: sp\_rodent:\*

12: sp\_virus:\*

13: sp\_vertebrate:\*

14: sp unclassified:\*

15: sp\_rvirus:\*

16: sp bacteriap:\*

17: sp\_archeap:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

용

Result Query

No. Score Match Length DB ID

Description

1	267	45.5	110	6	Q8HXV2	Q8hxv2 pongo pygma
2	251	42.8	110	6	Q8WNW6	Q8wnw6 felis silve
3	249	42.4	217	6	Q8WNE0	Q8wneO ateles geof
4	228	38.8	245	4	014644	Ol4644 homo sapien
5	219.5	37.4	106	13	Q9I8Q7	Q9i8q7 rana pipien
6	215	36.6	184	6	Q866T9	Q866t9 pan troglod
7	213	36.3	212	6	Q07368	Q07368 macaca mula
8	213	36.3	217	6	Q07367	Q07367 macaca mula
9	205	34.9	217	6	Q866U1	Q866ul pan troglod
10	201.5	34.3	110	13	Q98TA8	Q98ta8 pantodon bu
11	201	34.2	217	6	Q07369	Q07369 macaca mula
12	201	34.2	217	6	Q866T8	Q866t8 pan troglod
13	197	33.6	217	4	Q14407	Q14407 homo sapien
14	195.5	33.3	108	13	Q9DDE5	Q9dde5 brachydanio
15	195.5	33.3	108	13	Q90ZN4	Q90zn4 catla catla
16	195	33.2	111	13	Q98TB0	Q98tb0 chitala chi
17	195	33.2	217	6	Q8WND9	Q8wnd9 ateles geof
18	193.5	33.0	110	13	Q90ZY1	Q90zy1 hiodon alos
19	191.5	32.6	111	13	Q98TA7	Q98ta7 osteoglossu
20	189.5	32.3	108	13	Q98TB1	Q98tbl catostomus
21	187.5	31.9	87	13	Q98TA9	Q98ta9 gnathonemus
22	186	31.7	217	6	Q866U0	Q866u0 pan troglod
23	185.5	31.6	91	13	Q98TB2	Q98tb2 ambloplites
24	171.5	29.2	217	6	Q8MI74	Q8mi74 callithrix
25	161	27.4	216	11	070615	070615 spalax leuc
26	159.5	27.2	52	6	Q9TV91	Q9tv91 equus cabal
27	159.5	27.2	216	6	Q8MI73	Q8mi73 delphinus d
28	159.5	27.2	216	6	Q8HYE5	Q8hye5 ailuropoda
29	159.5	27.2	216	6	Q7YQB8	Q7yqb8 hippopotamu
30	156	26.6	216	11	Q9R2C3	Q9r2c3 mus musculu
31	155.5	26.5	216	6	Q7YRR6	Q7yrr6 camelus dro
32	154	26.2	216	11	Q9JKM4	Q9jkm4 cavia porce
33	152	25.9	178	6	Q95MJ5	Q95mj5 tarsius ban
34	149.5	25.5	204	6	Q95205	Q95205 ovis aries
35	149	25.4	132	13	Q8AV14	Q8av14 petromyzon
36	148.5	25.3	104	13	Q7T107	Q7t107 dicentrarch
37	148.5	25.3	108	13	0800И0	Q800n0 morone chry
38	148.5	25.3	108	13	Q800M9	Q800m9 morone saxa
39	148.5	25.3	108	13	0800M8	Q800m8 morone chry
40	148.5	25.3	108	13	Q800M7	Q800m7 morone amer
41	148.5	25.3	159	13	093607	093607 paralichthy
42	148.5	25.3	182	13	073720	073720 oreochromis
43	148.5	25.3	182	13	042289	042289 oreochromis
44	148.5	25.3	182	13	P79824	P79824 oreochromis
45	148.5	25.3	185	13	057436	057436 paralichthy
_						· · · · · · · · · · · · · · · · · · ·

#### ALIGNMENTS

```
RESULT 1
Q8HXV2
ID Q8HXV2 PRELIMINARY; PRT; 110 AA.
AC Q8HXV2;
DT 01-MAR-2003 (TrEMBLrel. 23, Created)
DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT 01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
```

```
DE
    Insulin precursor.
GN
    INS.
OS
    Pongo pygmaeus (Orangutan).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Pongo.
OC
OX
    NCBI TaxID=9600;
RN
     [1]
RP
    SEQUENCE FROM N.A.
RA
    Stead J.D.H., Jeffreys A.J.;
RT
    "Haplotype diversity at the insulin region.";
    Submitted (JUL-2002) to the EMBL/GenBank/DDBJ databases.
RL
    EMBL; AY137503; AAN06937.1; -.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
    GO; GO:0007582; P:physiological processes; IEA.
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
DR
    PRINTS; PR00277; INSULINB.
    SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
DR
    SEQUENCE 110 AA; 12038 MW; 22D2B32B94F520F8 CRC64;
SQ
 Query Match
                        45.5%; Score 267; DB 6; Length 110;
 Best Local Similarity 60.5%; Pred. No. 9e-24;
 Matches 52; Conservative
                              0; Mismatches 0; Indels 34; Gaps
          56 FVNOHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
             Db
          25 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRREAEDLQVGQVELGGGPGAGSLQPLALEG 84
          86 ----RGIVEQCCTSICSLYQLENYCN 107
Qу
                 11111111111
Dh
          85 SLQKRGIVEQCCTSICSLYQLENYCN 110
RESULT 2
O8MNM6
ΙD
              PRELIMINARY; PRT;
                                        110 AA.
    Q8WNW6
AC
    Q8WNW6;
    01-MAR-2002 (TrEMBLrel. 20, Created)
DT
    01-MAR-2002 (TrEMBLrel. 20, Last sequence update)
DT
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DE
    Preproinsulin.
OS
    Felis silvestris catus (Cat).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Carnivora; Fissipedia; Felidae; Felis.
OX
    NCBI TaxID=9685;
RN
    [1]
    SEQUENCE FROM N.A.
RP
RC
    TISSUE=Pancreas;
RA
    Okamoto S., Morimatsu M.;
RT
    "cat insulin.";
RL
    Submitted (MAY-2000) to the EMBL/GenBank/DDBJ databases.
CC
    -!- SUBCELLULAR LOCATION: SECRETED (BY SIMILARITY).
CC
    -!- SIMILARITY: BELONGS TO THE INSULIN/IGF/RELAXIN FAMILY.
DR
    EMBL; AB043535; BAB84110.1; -.
DR
    GO; GO:0005576; C:extracellular; IEA.
```

```
DR
    GO; GO:0007582; P:physiological processes; IEA.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
DR
    PROSITE; PS00262; INSULIN; 1.
SO
    SEQUENCE
             110 AA; 12069 MW; 95FB6E170C7BECA4 CRC64;
 Query Match
                        42.8%; Score 251; DB 6; Length 110;
 Best Local Similarity
                        55.8%; Pred. No. 7.1e-22;
          48; Conservative
                              2; Mismatches 2; Indels
 Matches
                                                             34; Gaps
                                                                         1;
          56 FVNOHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
             25 FVNQHLCGSHLVEALYLVCGERGFFYTPKARREAEDLQGKDAELGEAPGAGGLQPSALEA 84
Db
          86 ----RGIVEQCCTSICSLYQLENYCN 107
Οv
                 11111111111111111111
Db
          85 PLQKRGIVEQCCASVCSLYQLEHYCN 110
RESULT 3
Q8WNEO
                PRELIMINARY;
                                 PRT;
                                        217 AA.
TD
    Q8WNE0
AC
    O8WNEO;
    01-MAR-2002 (TrEMBLrel. 20, Created)
DT
    01-MAR-2002 (TrEMBLrel. 20, Last sequence update)
TП
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
    Growth hormone.
    GH-N.
GN
OS
    Ateles geoffroyi (Black-handed spider monkey).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Primates; Platyrrhini; Cebidae; Atelinae; Ateles.
OC
OX
    NCBI TaxID=9509;
RN
    [1]
RP
    SEQUENCE FROM N.A.
    Revol A., Esquivel D., Santiago D., Barrera-Saldana H.;
RA
RT
    "Independent duplication of the growth hormone gene in three
RT
    Anthropoidean lineages.";
RL
    Submitted (APR-2001) to the EMBL/GenBank/DDBJ databases.
DR
    EMBL; AF374234; AAL72286.1; -.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
    InterPro; IPR001400; Somatotropin.
    Pfam; PF00103; hormone; 1.
DR
DR
    PRINTS; PR00836; SOMATOTROPIN.
DR
    PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
    PROSITE; PS00338; SOMATOTROPIN 2; 1.
SQ
    SEQUENCE 217 AA; 24894 MW; 425829FF41EEAAE6 CRC64;
 Query Match
                        42.4%; Score 249; DB 6; Length 217;
 Best Local Similarity 97.9%; Pred. No. 2.7e-21;
 Matches
           47; Conservative
                               0; Mismatches
                                               1; Indels
                                                              0; Gaps
                                                                         0;
Qу
           2 FPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
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DR

GO; GO:0005179; F:hormone activity; IEA.

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RESULT 4
014644
ΙD
    014644
                PRELIMINARY;
                                  PRT;
                                         245 AA.
    014644;
AC
DT
    01-JAN-1998 (TrEMBLrel. 05, Created)
    01-JAN-1998 (TrEMBLrel. 05, Last sequence update)
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
    Placental growth hormone isoform hGH-V3 precursor.
DE
    HGH-V.
GN
OS
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
    NCBI TaxID=9606;
OX
RN
    [1]
RP
    SEQUENCE FROM N.A.
RC
    TISSUE=Term placenta;
    MEDLINE=98373737; PubMed=9709963;
RX
RA
    Boguszewski C.L., Svensson P.A., Jansson T., Clark R.,
RA
    Carlsson L.M.S., Carlsson B.;
    "Cloning of two novel growth hormone transcripts expressed in human
RT
    placenta.";
RT
    J. Clin. Endocrinol. Metab. 83:2878-2885(1998).
RL
    EMBL; AF006061; AAB71829.1; -.
DR
    HSSP; P01241; 1A22.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
    InterPro; IPR001400; Somatotropin.
    Pfam; PF00103; hormone; 1.
DR
    PRINTS; PR00836; SOMATOTROPIN.
DR
    PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
KW
    Signal.
FT
    SIGNAL
                  1
                        26
                                 POTENTIAL.
    SEQUENCE
               245 AA; 27101 MW; 14CC7F8CD75D91C8 CRC64;
SQ
                         38.8%; Score 228; DB 4; Length 245;
  Query Match
  Best Local Similarity 91.7%; Pred. No. 9.5e-19;
                                2; Mismatches 2; Indels
                                                                0; Gaps
                                                                            0;
  Matches
          44; Conservative
           2 FPTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
Qv
             27 FPTIPLSRLFDNAMLRARRLYQLAYDTYQEFEEAYILKEQKYSFLQNP 74
Db
RESULT 5
Q9I8Q7
    Q918Q7
                PRELIMINARY;
                                  PRT;
                                         106 AA.
ID
    Q9I8Q7;
     01-OCT-2000 (TrEMBLrel. 15, Created)
DT
DT
     01-OCT-2000 (TrEMBLrel. 15, Last sequence update)
     01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
    Preproinsulin.
OS
    Rana pipiens (Northern leopard frog).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Amphibia; Batrachia; Anura; Neobatrachia; Ranoidea; Ranidae; Rana.
OC.
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OX
    NCBI TaxID=8404;
RN
    [1]
RP
    SEQUENCE FROM N.A.
RX
    MEDLINE=20362507; PubMed=10818274;
RA
    Irwin D.M., Sivarajah P.;
     "Proinsulin cDNAs from the leopard frog, Rana pipiens: evolution of
RT
RT
    proinsulin processing.";
    Comp. Biochem. Physiol. 125B:405-410(2000).
RL
CC
    -!- SUBCELLULAR LOCATION: SECRETED (BY SIMILARITY).
    -!- SIMILARITY: BELONGS TO THE INSULIN/IGF/RELAXIN FAMILY.
CC
    EMBL; AF227187; AAF87285.1; -.
DR
    HSSP; P01315; 1SDB.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
DR
    GO; GO:0005179; F:hormone activity; IEA.
    GO; GO:0007582; P:physiological processes; IEA.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
DR
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
DR
    SEQUENCE 106 AA; 12183 MW; 3A870EEC70217F92 CRC64;
SQ
                         37.4%; Score 219.5; DB 13; Length 106;
 Query Match
 Best Local Similarity 49.4%; Pred. No. 3.7e-18;
          41; Conservative
                               7; Mismatches 4; Indels
                                                               31; Gaps
 Matches
                                                                           1;
          56 FVNOHLCGSHLVEALYLVCGERGFFYTPKTR----- 86
Qу
             24 FDNQYLCGSHLVEALYMVCGDRGFFYSPRSRRDLEQPLVNGLQGSELDEMQVQSQAFQKR 83
Db
          87 -- GIVEQCCTSICSLYQLENYCN 107
Qу
               1111111 : 1111 111111
Db
          84 KPGIVEQCCHNTCSLYDLENYCN 106
RESULT 6
Q866T9
                PRELIMINARY;
                                  PRT;
                                         184 AA.
ID
    Q866T9
AC
     Q866T9;
DT
     01-JUN-2003 (TrEMBLrel. 24, Created)
DT
     01-JUN-2003 (TrEMBLrel. 24, Last sequence update)
DT
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DE
     Placental lactogen PL-C (Fragment).
     Pan troglodytes (Chimpanzee).
OS
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Pan.
OC
OX
    NCBI TaxID=9598;
RN
     [1]
RP
     SEQUENCE FROM N.A.
RA
     Revol A., Esquivel D.E., Barrera H.S.;
RT
     "The GH-PL locus a hot-point between human and chimpanzee genomes.";
     Submitted (AUG-2002) to the EMBL/GenBank/DDBJ databases.
RL
DR
    EMBL; AY146627; AAN84507.1; -.
DR
     GO; GO:0005576; C:extracellular; IEA.
DR
     GO; GO:0005179; F:hormone activity; IEA.
     InterPro; IPR001400; Somatotropin.
DR
DR
     Pfam; PF00103; hormone; 1.
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E

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DR
    PRINTS; PR00836; SOMATOTROPIN.
DR
    PROSITE; PS00266; SOMATOTROPIN 1; 1.
FT
    NON TER
                184
                       184
    SEQUENCE
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SQ
                         36.6%; Score 215; DB 6; Length 184;
 Query Match
 Best Local Similarity
                         85.1%; Pred. No. 2.4e-17;
                               4; Mismatches
                                                                          0;
 Matches
           40; Conservative
                                                 3; Indels
                                                                  Gaps
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Qу
             27 FPTIPLSRLFDHAMLQAHRAHQLAIDTYQEFEEAYIPKDQKYSFLHD 73
Db
RESULT 7
Q07368
ID
    007368
                PRELIMINARY;
                                 PRT:
                                        212 AA.
AC
    007368;
DT
    01-NOV-1996 (TrEMBLrel. 01, Created)
    01-NOV-1996 (TrEMBLrel. 01, Last sequence update)
DT
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
    Somatotropin 2 precursor (Growth hormone 2) (Fragment).
    Macaca mulatta (Rhesus macaque).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
    Mammalia; Eutheria; Primates; Catarrhini; Cercopithecidae;
OC
    Cercopithecinae; Macaca.
OX
    NCBI TaxID=9544;
RN
    [1]
RP
    SEQUENCE FROM N.A.
    TISSUE=Placenta;
RC
    MEDLINE=94008724; PubMed=8404617;
RX
    Golos T.G., Durning M., Fisher J.M., Fowler P.D.;
RA
RT
    "Cloning of four growth hormone/chorionic somatomammotropin-related
RT
     complementary deoxyribonucleic acids differentially expressed during
RT
    pregnancy in the rhesus monkey placenta.";
    Endocrinology 133:1744-1752(1993).
RL
    EMBL; L16553; AAA18840.1; -.
DR
DR
    PIR; 167408; 167408.
    HSSP; P01241; 1AXI.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
DR
    InterPro; IPR001400; Somatotropin.
    Pfam; PF00103; hormone; 1.
DR
    PRINTS; PR00836; SOMATOTROPIN.
DR
    PROSITE; PS00338; SOMATOTROPIN 2; 1.
DR
FT
    NON TER
                  1
    SEQUENCE
               212 AA;
                      24525 MW; 27BC91106256E6F5 CRC64;
SQ
  Query Match
                         36.3%; Score 213; DB 6;
                                                   Length 212;
  Best Local Similarity
                         78.7%; Pred. No. 4.8e-17;
  Matches
           37; Conservative
                               9; Mismatches
                                                 1; Indels
                                                               0;
                                                                   Gaps
                                                                          0;
Qy
           3 PTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
             Db
          23 PSVPLSRLFDHAMIQAHRLHQLAFDTYQEFEEAYIPKEKKHSLMENP 69
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RESULT 8
007367
ID
     Q07367
                 PRELIMINARY;
                                   PRT;
                                         217 AA.
     Q07367;
DT
     01-NOV-1996 (TrEMBLrel. 01, Created)
DT
     01-NOV-1996 (TrEMBLrel. 01, Last sequence update)
     01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
     Chorionic somatomammotropin-1.
OS
    Macaca mulatta (Rhesus macaque).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Cercopithecidae;
OC
     Cercopithecinae; Macaca.
OX
    NCBI TaxID=9544;
RN
     [1]
RP
     SEOUENCE FROM N.A.
RC
    TISSUE=Midpregnancy placenta;
    MEDLINE=94008724; PubMed=8404617;
RX
    Golos T.G., Durning M., Fisher J.M., Fowler P.D.;
RA
     "Cloning of four growth hormone/chorionic somatomammotropin-related
RT
     complementary deoxyribonucleic acids differentially expressed during
RT
     pregnancy in the rhesus monkey placenta.";
RT
     Endocrinology 133:1744-1752(1993).
RL
     EMBL; L16552; AAA18839.1; -.
DR
DR
     PIR; I53267; I53267.
DR
    HSSP; P01241; 1AXI.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
     GO; GO:0005179; F:hormone activity; IEA.
DR
     InterPro; IPR001400; Somatotropin.
DR
     Pfam; PF00103; hormone; 1.
DR
     PRINTS; PR00836; SOMATOTROPIN.
DR
     PROSITE; PS00338; SOMATOTROPIN 2; 1.
SQ
               217 AA; 24942 MW; FF5AA8915131F2BC CRC64;
  Query Match
                          36.3%; Score 213; DB 6; Length 217;
  Best Local Similarity
                         78.7%; Pred. No. 5e-17;
            37; Conservative
                                9; Mismatches
                                                  1;
                                                      Indels
                                                                    Gaps
                                                                             0;
Qу
            3 PTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
              Db
           28 PSVPLSRLFDHAMIQAHRLHQLAFDTYQEFEEAYIPKEKKHSLMENP 74
RESULT 9
Q866U1
ID
     0866U1
                 PRELIMINARY;
                                  PRT;
                                         217 AA.
AC
     Q866U1;
DT
     01-JUN-2003 (TrEMBLrel. 24, Created)
     01-JUN-2003 (TrEMBLrel. 24, Last sequence update)
DΤ
DΤ
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DE
     Placental lactogen PL-A.
OS
     Pan troglodytes (Chimpanzee).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Pan.
OC
OX
     NCBI TaxID=9598;
RN
     [1]
RP
     SEQUENCE FROM N.A.
RA
     Revol A., Esquivel D.E., Barrera H.S.;
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"The GH-PL locus a hot-point between human and chimpanzee genomes.";
     Submitted (AUG-2002) to the EMBL/GenBank/DDBJ databases.
RL
DR
     EMBL; AY146625; AAN84505.1; -.
DR
     GO; GO:0005576; C:extracellular; IEA.
     GO; GO:0005179; F:hormone activity; IEA.
DR
     InterPro; IPR001400; Somatotropin.
DR
DR
     Pfam; PF00103; hormone; 1.
     PRINTS; PR00836; SOMATOTROPIN.
DR
     PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
     PROSITE; PS00338; SOMATOTROPIN 2; 1.
DR
               217 AA; 25081 MW; C74B6262D8A93060 CRC64;
     SEQUENCE
SQ
                          34.9%; Score 205; DB 6; Length 217;
  Query Match
  Best Local Similarity
                         84.8%; Pred. No. 4.4e-16;
                                4: Mismatches
                                                      Indels
                                                                0; Gaps
                                                                            0;
           39; Conservative
                                                  3;
  Matches
           4 TIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
Qу
              29 TVPLSRLFDHAMLQAHRAYQLAIDTYQEFEEAYILKEQKYSFLQNP 74
Db
RESULT 10
098TA8
                PRELIMINARY;
                                  PRT;
                                         110 AA.
     Q98TA8
TD
AC
     Q98TA8;
     01-JUN-2001 (TrEMBLrel. 17, Created)
     01-JUN-2001 (TrEMBLrel. 17, Last sequence update)
DΤ
     01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
     Preproinsulin.
     Pantodon buchholtzi (Butterflyfish).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Actinopterygii; Neopterygii; Teleostei; Osteoglossomorpha;
OC
     Osteoglossiformes; Pantodontidae; Pantodon.
OC
OX
     NCBI TaxID=8276;
RN
     [1]
RP
     SEQUENCE FROM N.A.
     MEDLINE=21203577; PubMed=11306171;
RX
     Al-Mahrouki A.A., Irwin D.M., Graham L.C., Youson J.H.;
RA
     "Molecular cloning of preproinsulin cDNAs from several
RT
RT
     osteoglossomorphs and a cyprinid.";
RL
     Mol. Cell. Endocrinol. 174:51-58(2001).
     -!- SUBCELLULAR LOCATION: SECRETED (BY SIMILARITY).
CC
CC
     -!- SIMILARITY: BELONGS TO THE INSULIN/IGF/RELAXIN FAMILY.
     EMBL; AF199588; AAK28712.1; -.
DR
     HSSP; P01308; 1HIS.
DR
     GO; GO:0005576; C:extracellular; IEA.
DR
     GO; GO:0005179; F:hormone activity; IEA.
DR
DR
     GO; GO:0007582; P:physiological processes; IEA.
DR
     InterPro; IPR004825; Ins/IGF/relax.
DR
     Pfam; PF00049; Insulin; 1.
DR
     PRINTS; PR00277; INSULINB.
DR
     SMART; SM00078; IlGF; 1.
DR
     PROSITE; PS00262; INSULIN; 1.
SQ
     SEQUENCE
                110 AA; 12324 MW; BDECCD659D872E06 CRC64;
                          34.3%; Score 201.5; DB 13; Length 110;
  Query Match
  Best Local Similarity 43.5%; Pred. No. 5.2e-16;
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RТ

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5; Indels
                                                            35; Gaps
 Matches
           37; Conservative
                               8: Mismatches
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Qy
             26 SQHLCGSHLVDALYMVCGEKGFFYQPKTKRDVDPLLGFLSPKSAQENEADEYPYKDQGDL 85
Db
          86 --- RGIVEQCCTSICSLYQLENYCN 107
Qу
                86 KVKRGIVEQCCHHPCNIFDLQNYCN 110
Db
RESULT 11
007369
                PRELIMINARY;
                                PRT;
                                       217 AA.
ID
    007369
    007369;
AC
    01-NOV-1996 (TrEMBLrel. 01, Created)
DТ
    01-NOV-1996 (TrEMBLrel. 01, Last sequence update)
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
    Chorionic somatomammotropin-3.
DΕ
    Macaca mulatta (Rhesus macaque).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Cercopithecidae;
OC
    Cercopithecinae; Macaca.
OC
    NCBI TaxID=9544;
OX
RN
    SEQUENCE FROM N.A.
RP
RC
    TISSUE=Midpregnancy placenta;
    MEDLINE=94008724; PubMed=8404617;
RX
    Golos T.G., Durning M., Fisher J.M., Fowler P.D.;
RA
    "Cloning of four growth hormone/chorionic somatomammotropin-related
RТ
    complementary deoxyribonucleic acids differentially expressed during
RT
RT
    pregnancy in the rhesus monkey placenta.";
    Endocrinology 133:1744-1752(1993).
RI_1
DR
    EMBL; L16554; AAA18841.1; -.
    PIR; 167409; 167409.
DR
    HSSP; P01241; 1AXI.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
    InterPro; IPR001400; Somatotropin.
DR
    Pfam; PF00103; hormone; 1.
DR
    PRINTS; PRO0836; SOMATOTROPIN.
DR
    PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
    PROSITE; PS00338; SOMATOTROPIN 2; 1.
DR
    SEOUENCE
               217 AA; 24874 MW; F1EB6AFDBBA1B185 CRC64;
SQ
                        34.2%; Score 201; DB 6; Length 217;
  Query Match
  Best Local Similarity
                        74.5%; Pred. No. 1.3e-15;
  Matches
           35; Conservative
                              8: Mismatches
                                               4; Indels
                                                             0; Gaps
                                                                         0;
           3 PTIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQNP 49
Qу
             Dh
          28 PSVPLSRLFDNIMMQAHRLHQLAFDTYQEFEKTYIPKEKKHSLMGNP 74
RESULT 12
Q866T8
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                                       217 AA.
ID Q866T8
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AC
     Q866T8;
DT
     01-JUN-2003 (TrEMBLrel. 24, Created)
DT
     01-JUN-2003 (TrEMBLrel. 24, Last sequence update)
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DE
     Placental lactogen PL-D.
OS
     Pan troglodytes (Chimpanzee).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Pan.
OC
    NCBI_TaxID=9598;
OX
RN
     [1]
     SEQUENCE FROM N.A.
RP
     Revol A., Esquivel D.E., Barrera H.S.;
RA
     "The GH-PL locus a hot-point between human and chimpanzee genomes.";
RT
     Submitted (AUG-2002) to the EMBL/GenBank/DDBJ databases.
RL
    EMBL; AY146628; AAN84508.1; -.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
     InterPro; IPR001400; Somatotropin.
     Pfam; PF00103; hormone; 1.
DR
     PRINTS; PR00836; SOMATOTROPIN.
DR
     PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
     PROSITE; PS00338; SOMATOTROPIN 2; 1.
DR
SQ
     SEQUENCE
               217 AA; 25135 MW; 1EB7B89B8A12E4F4 CRC64;
                         34.2%; Score 201; DB 6; Length 217;
  Query Match
  Best Local Similarity
                         82.2%; Pred. No. 1.3e-15;
                                5; Mismatches
                                                  3; Indels
                                                                0; Gaps
                                                                            0;
  Matches
           37; Conservative
            4 TIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQN 48
Qу
              29 TVPLSRLFDHAMLQAHRAHQLAIDTYQEFEEAYIPKDQKYSFLHD 73
RESULT 13
Q14407
                PRELIMINARY;
                                  PRT;
                                         217 AA.
ID
     Q14407
AC
     Q14407;
     01-NOV-1996 (TrEMBLrel. 01, Created)
DT
     01-NOV-1996 (TrEMBLrel. 01, Last sequence update)
DT
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DE
     Chorionic somatomammotropin CS-2 (Chorionic somatomammotropin hormone
DE
     2).
OS
     Homo sapiens (Human).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
     NCBI TaxID=9606;
RN
     [1]
RP
     SEQUENCE FROM N.A.
RX
     MEDLINE=89307277; PubMed=2744760;
RA
     Chen E.Y., Liao Y.C., Smith D.H., Barrera-Saldana H.A., Gelinas R.E.,
RA
     Seeburg P.H.;
RT
     "The human growth hormone locus: nucleotide sequence, biology, and
RT
     evolution.";
RL
     Genomics 4:479-497(1989).
RN
     [2]
RP
     SEQUENCE FROM N.A.
RX
     MEDLINE=91102558; PubMed=1980158;
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RA
    Vnencak-Jones C.L., Phillips J.A. III.;
    "Hot spots for growth hormone gene deletions in homologous regions
RT
RT
    outside of Alu repeats.";
    Science 250:1745-1748(1990).
RL
RN
     [3]
RP
    SEQUENCE FROM N.A.
    TISSUE=Placenta;
RC
RA
    Strausberg R.;
     Submitted (JUL-2002) to the EMBL/GenBank/DDBJ databases.
RL
DR
    EMBL; J03071; AAA52553.1; -.
DR
    EMBL; BC022044; AAH22044.1; -.
    EMBL; BC035965; AAH35965.1; -.
DR
    PIR; E32435; E32435.
DR
    HSSP; P01241; 1A22.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
DR
    InterPro; IPR001400; Somatotropin.
DR
    Pfam; PF00103; hormone; 1.
DR
    PRINTS; PR00836; SOMATOTROPIN.
DR
     PROSITE; PS00266; SOMATOTROPIN 1; 1.
DR
     PROSITE; PS00338; SOMATOTROPIN 2; 1.
DR
SO
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                         33.6%; Score 197; DB 4; Length 217;
  Query Match
  Best Local Similarity
                         80.0%; Pred. No. 3.9e-15;
                                                                0; Gaps
                                                                           0;
          36; Conservative
                                5; Mismatches
                                                 4; Indels
           4 TIPLSRLFDNAMLRAHRLHQLAFDTYQEFEEAYIPKEQKYSFLQN 48
Qу
              29 TVPLSRLFDHAMLQAHRAHQLAIDTYQEFEETYIPKDQKYSFLHD 73
Db
RESULT 14
Q9DDE5
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                                  PRT; 108 AA.
ID
     Q9DDE5
AC
     Q9DDE5;
     01-MAR-2001 (TrEMBLrel. 16, Created)
DT
     01-MAR-2001 (TrEMBLrel. 16, Last sequence update)
DT
DT
     01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DE
     Insulin precursor.
GN
     INS.
     Brachydanio rerio (Zebrafish) (Danio rerio).
OS
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Actinopterygii; Neopterygii; Teleostei; Ostariophysi; Cypriniformes;
OC
     Cyprinidae; Danio.
OX
     NCBI TaxID=7955;
RN
     [1]
RP
     SEQUENCE FROM N.A.
RX
     MEDLINE=99425190; PubMed=10495291;
     Argenton F., Zecchin E., Bortolussi M.;
RT
     "Early appearance of pancreatic hormone-expressing cells in the
RT
     zebrafish embryo.";
RL
     Mech. Dev. 87:217-221(1999).
CC
     -!- SUBCELLULAR LOCATION: SECRETED (BY SIMILARITY).
CC
     -!- SIMILARITY: BELONGS TO THE INSULIN/IGF/RELAXIN FAMILY.
DR
     EMBL; AJ237750; CAC20109.1; -.
DR
     HSSP; P01308; 1LPH.
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ZFIN; ZDB-GENE-980526-110; ins.
DR
    GO; GO:0005576; C:extracellular; IEA.
DR
    GO; GO:0005179; F:hormone activity; IEA.
    GO; GO:0007582; P:physiological processes; IEA.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
DR
    Pfam; PF00049; Insulin; 1.
    PRINTS; PR00277; INSULINB.
DR
DR
    SMART; SM00078; IlGF; 1.
    PROSITE; PS00262; INSULIN; 1.
DR
KW
    Signal.
                       23
    SIGNAL
                 1
                                POTENTIAL.
FT
                 24
                       53
                                INSULIN B CHAIN.
FT
    CHAIN
                 86
                     108
                                INSULIN A CHAIN.
FT
    CHAIN
    SEQUENCE 108 AA; 11904 MW; 3195289E72AD6D25 CRC64;
SO
                        33.3%; Score 195.5; DB 13; Length 108;
 Query Match
 Best Local Similarity 45.1%; Pred. No. 2.6e-15;
                               5; Mismatches
 Matches
          37; Conservative
                                                7; Indels
                                                             33; Gaps
          59 QHLCGSHLVEALYLVCGERGFFYTPK-----T 85
Qу
             27 QHLCGSHLVDALYLVCGPTGFFYNPKRDVEPLLGFLPPKSAQETEVADFAFKDHAELIRK 86
          86 RGIVEQCCTSICSLYQLENYCN 107
Qу
             87 RGIVEQCCHKPCSIFELQNYCN 108
Db
RESULT 15
Q90ZN4
ID
    Q90ZN4
                PRELIMINARY;
                                 PRT;
                                        108 AA.
AC
    Q90ZN4;
    01-DEC-2001 (TrEMBLrel. 19, Created)
    01-DEC-2001 (TrEMBLrel. 19, Last sequence update)
    01-JUN-2003 (TrEMBLrel. 24, Last annotation update)
DT
DE
    Preproinsulin.
    Catla catla (catla).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Actinopterygii; Neopterygii; Teleostei; Ostariophysi; Cypriniformes;
OC
OC
    Cyprinidae; Catla.
    NCBI TaxID=72446;
OX
RN
     SEOUENCE FROM N.A.
RP
     Bhattacharya S., Roy S.S., Dasgupta S., Ravikumar L., Mukherjee M.,
RA
     Bandyopadhyaya I., Wakabayasi K.;
RA
RT
     "A new cell secreting insulin.";
     Submitted (APR-2001) to the EMBL/GenBank/DDBJ databases.
RL
CC
     -!- SUBCELLULAR LOCATION: SECRETED (BY SIMILARITY).
CC
     -!- SIMILARITY: BELONGS TO THE INSULIN/IGF/RELAXIN FAMILY.
     EMBL; AF373021; AAK51558.1; -.
DR
     HSSP; P01308; 1LNP.
DR
     GO; GO:0005576; C:extracellular; IEA.
     GO; GO:0005179; F:hormone activity; IEA.
DR
DR
     GO; GO:0007582; P:physiological processes; IEA.
DR
     InterPro; IPR004825; Ins/IGF/relax.
DR
     Pfam; PF00049; Insulin; 1.
DR
     PRINTS; PR00277; INSULINB.
```

```
SMART; SM00078; IlGF; 1.
DR
DR PROSITE; PS00262; INSULIN; 1.
    SEQUENCE 108 AA; 11881 MW; D713026E22EF5D59 CRC64;
SQ
 Query Match 33.3%; Score 195.5; DB 13; Length 108; Best Local Similarity 45.1%; Pred. No. 2.6e-15; Matches 37; Conservative 5; Mismatches 7; Indels 33; Gaps 1;
           59 QHLCGSHLVEALYLVCGERGFFYTPK----T 85
Qу
             Db
           27 QHLCGSHLVDALYLVCGPTGFFYNPKRDVDPLMGFLPPKSAQETEVADFAFKDHAEVIRK 86
          86 RGIVEQCCTSICSLYQLENYCN 107
Qу
             87 RGIVEQCCHKPCSIFELQNYCN 108
Db
```

Search completed: July 15, 2004, 16:41:00 Job time: 36.6984 secs

## GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: July 15, 2004, 16:28:49; Search time 7.38619 Seconds

(without alignments)

754.314 Million cell updates/sec

Title: US-09-423-100-6

Perfect score: 587

Sequence: 1 MFPTIPLSRLFDNAMLRAHR.....IVEQCCTSICSLYQLENYCN 107

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 141681 seqs, 52070155 residues

Total number of hits satisfying chosen parameters: 141681

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database : SwissProt\_42:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

		8				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
1	273.5	46.6	51	1	INS BALPH	P01312 balaenopter
2	273.5	46.6	51	1	INS ELEMA	P01316 elephas max
3	273	46.5	110	1	INS CERAE	P30407 cercopithec
4	273	46.5	110	1	INS MACFA	P30406 macaca fasc
5	268.5	45.7	51	1	INS ACOCA	P01324 acomys cahi
6	267	45.5	110	1	INS HUMAN	P01308 homo sapien
7	267	45.5	110	1	INS PANTR	P30410 pan troglod
8	266	45.3	110	1	INS_SPETR	Q91xi3 spermophilu
9	263.5	44.9	51	1	INS_BALBO	P01314 balaenopter
10	263.5	44.9	51	1	INS_CAMDR	P01320 camelus dro
11	263.5	44.9	51	1	INS CAPHI	P01319 capra hircu
12	263	44.8	108	1	INS PIG	P01315 sus scrofa
13	263	44.8	110	1	INS RABIT	P01311 oryctolagus
14	262.5	44.7	51	1	INS FELCA	P06306 felis silve
15	262	44.6	110	1	INS CANFA	P01321 canis famil
16	260	44.3	110	1	INS CRILO	P01313 cricetulus
17	258.5	44.0	105	1	INS_BOVIN	P01317 bos taurus

18	257	43.8	108	1	INS AOTTR	P10604 aotus trivi
19	257	43.8	110	1	INS PSAOB	Q62587 psammomys o
20	256.5	43.7	51	1	INS DIDMA	P18109 didelphis m
21	255.5	43.5	217	1	SOMA HUMAN	P01241 homo sapien
22	255.5	43.5	217	1	SOMA MACMU	P33093 macaca mula
23	255.5	43.5	217	1	SOMA PANTR	P58756 pan troglod
24	254.5	43.4	105	1	INS_SHEEP	P01318 ovis aries
25	252	42.9	86	1	INS HORSE	P01310 equus cabal
26	251.5	42.8	51	1	INS CHIBR	P01327 chinchilla
27	250	42.6	108	1	INST MOUSE	P01325 mus musculu
28	249	42.4	110	1	INS1 RAT	P01322 rattus norv
29	249	42.4	217	1	SOMA CALJA	Q9gmb3 callithrix
30	249	42.4	217	1	SOMA_SAIBB	P58343 saimiri bol
31	248.5	42.3	51	1	INS ANSAN	P07454 anser anser
32	248	42.2	110	1	INSZ MOUSE	P01326 mus musculu
33	248	42.2	110	1	INS2 RAT	P01323 rattus norv
34	246	41.9	52	1	INS_ACIGU	P81423 acipenser g
35	245	41.7	103	1	INS_SELRF	P51463 selasphorus
36	244.5	41.7	51	1	INS_HYSCR	P01328 hystrix cri
37	244.5	41.7	51	1.	INS_TRASC	P31887 trachemys s
38	239.5	40.8	107	1	INS_CHICK	P01332 gallus gall
39	236	40.2	217	1	SOM2_PANTR	P58757 pan troglod
40	235.5	40.1	51	1	INS_ORNAN	Q9tqy7 ornithorhyn
41	233.5	39.8	81	1	INS_ANAPL	P01333 anas platyr
42	231.5	39.4	51	1	INS_ALLMI	P12703 alligator m
43	231	39.4	52	1	INS_LEPSP	P09476 lepisosteus
44	228.5	38.9	51	1	INS_ZAODH	P12708 zaocys dhum
45	228	38.8	217	1	SOM2_HUMAN	P01242 homo sapien

## ALIGNMENTS

```
RESULT 1
INS BALPH
     INS BALPH
                    STANDARD;
                                    PRT;
                                            51 AA.
ID
AC
     P01312;
     21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
     Insulin.
GN
     INS.
     Balaenoptera physalus (Finback whale) (Common rorqual), and
OS
     Physeter catodon (Sperm whale) (Physeter macrocephalus).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Cetartiodactyla; Cetacea; Mysticeti;
OC
OC
     Balaenopteridae; Balaenoptera.
OX
     NCBI_TaxID=9770, 9755;
RN
RP
     PARTIAL SEQUENCE.
RC
     SPECIES=B.physalus;
RA
     Hama H., Titani K., Sakaki S., Narita K.;
RT
     "The amino acid sequence in fin-whale insulin.";
     J. Biochem. 56:285-293(1964).
RL
RN
     [2]
RP
     SEQUENCE.
RC
     SPECIES=P.catodon;
```

```
RA
     Ishihara Y., Saito T., Ito Y., Fujino M.;
RT
     "Structure of sperm- and sei-whale insulins and their breakdown by
RT
     whale pepsin.";
     Nature 181:1468-1469(1958).
RL
RN
     [3]
RP
     SEQUENCE.
RC
     SPECIES=P.catodon;
RA
     Harris J.I., Sanger F., Naughton M.A.;
RT
     "Species differences in insulin.";
     Arch. Biochem. Biophys. 65:427-438(1956).
RL
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
         increases cell permeability to monosaccharides, amino acids and
CC
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
         cycle, and glycogen synthesis in liver.
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
         disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
CC
     -!- SIMILARITY: Belongs to the insulin family.
CC
DR
     PIR; A91918; INWHF.
     PIR; A93142; INWHP.
DR
     HSSP; P01317; 1APH.
DR
     InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     PRINTS; PR00277; INSULINB.
DR
     SMART; SM00078; IlGF; 1.
DR
     PROSITE; PS00262; INSULIN; 1.
     Insulin family; Hormone; Glucose metabolism.
KW
FT
                         30
                                  INSULIN B CHAIN.
     CHAIN
                  1
FT
     NON CONS
                  30
                         31
FT
     CHAIN
                  31
                         51
                                  INSULIN A CHAIN.
                  7
                         37
FT
     DISULFID
                                  INTERCHAIN.
FT
     DISULFID
                  19
                         50
                                  INTERCHAIN.
FT
     DISULFID
                  36
                         41
     SEQUENCE
                51 AA; 5766 MW; 9007B514691A7CDD CRC64;
SO
  Query Match
                          46.6%;
                                  Score 273.5; DB 1;
                                                       Length 51;
  Best Local Similarity
                          96.2%;
                                  Pred. No. 1.9e-22;
  Matches
            50; Conservative
                                 0; Mismatches
                                                   1; Indels
                                                                 1; Gaps
                                                                             1;
           56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              Db
            1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCTSICSLYQLENYCN 51
RESULT 2
INS ELEMA
     INS ELEMA
                    STANDARD;
                                   PRT;
                                           51 AA.
TD
AC
     P01316;
DT
     21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
     10-OCT-2003 (Rel. 42, Last annotation update)
DТ
DE
     Insulin.
GN
     INS.
OS
     Elephas maximus (Indian elephant).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Proboscidea; Elephantidae; Elephas.
OX
     NCBI TaxID=9783;
RN
     [1]
```

```
RP
    SEOUENCE.
RX
    MEDLINE=66160119; PubMed=5949593;
RA
    Smith L.F.;
RT
     "Species variation in the amino acid sequence of insulin.";
RL
     Am. J. Med. 40:662-666(1966).
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
         increases cell permeability to monosaccharides, amino acids and
CC
CC
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
         cycle, and glycogen synthesis in liver.
CC
    -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
    -!- SUBCELLULAR LOCATION: Secreted.
CC
    -!- MISCELLANEOUS: THE SPECIES OF ELEPHANT IS NOT GIVEN, BUT IT IS
CC
        MOST PROBABLY THE INDIAN ELEPHANT (ELEPHAS MAXIMUS).
CC
CC
    -!- SIMILARITY: Belongs to the insulin family.
DR
    HSSP; P01308; 1AIO.
DR
    InterPro; IPR004825; Ins/IGF/relax.
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
DR
    Insulin family; Hormone; Glucose metabolism.
KW
FT
    CHAIN
                  1
                        30
                                 INSULIN B CHAIN.
FT
    NON CONS
                 30
                        31
FT
    CHAIN
                 31
                        51
                                 INSULIN A CHAIN.
                 7
                        37
FT
    DISULFID
                                 INTERCHAIN.
FT
    DISULFID
                 19
                        50
                                 INTERCHAIN.
FT
    DISULFID
                 36
                       41
SO
    SEQUENCE 51 AA; 5752 MW; 9007B50CDB457D6D CRC64;
  Query Match
                         46.6%; Score 273.5; DB 1; Length 51;
  Best Local Similarity
                         94.2%; Pred. No. 1.9e-22;
           49; Conservative
                                1; Mismatches
                                                  1; Indels
                                                                1; Gaps
                                                                            1;
          56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              1 FVNQHLCGSHLVEALYLVCGERGFFYTPKT-GIVEQCCTGVCSLYQLENYCN 51
RESULT 3
INS CERAE
     INS CERAE
ΙD
                   STANDARD;
                                  PRT;
                                         110 AA.
     P30407; P01309;
AC
DT
     01-APR-1993 (Rel. 25, Created)
DT
     01-APR-1993 (Rel. 25, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DΕ
    Insulin precursor.
GN
     INS.
OS
     Cercopithecus aethiops (Green monkey) (Grivet).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
    Mammalia; Eutheria; Primates; Catarrhini; Cercopithecidae;
OC
     Cercopithecinae; Cercopithecus.
     NCBI TaxID=9534;
OX
RN
     [1]
RP
     SEQUENCE FROM N.A.
RX
    MEDLINE=92219953; PubMed=1560757;
RA
     Seino S., Bell G.I., Li W.;
RT
     "Sequences of primate insulin genes support the hypothesis of a
```

```
RT
    slower rate of molecular evolution in humans and apes than in
RT
    monkeys.";
    Mol. Biol. Evol. 9:193-203(1992).
RL
RN
    SEQUENCE OF 57-87.
RP
    MEDLINE=72258016; PubMed=4626369;
RX
    Peterson J.D., Nehrlich S., Oyer P.E., Steiner D.F.;
RA
    "Determination of the amino acid sequence of the monkey, sheep, and
RT
    dog proinsulin C-peptides by a semi-micro Edman degradation
RT
RT
    procedure.";
    J. Biol. Chem. 247:4866-4871(1972).
RL
    -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
        increases cell permeability to monosaccharides, amino acids and
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
CC
    -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
       disulfide bonds.
    -!- SUBCELLULAR LOCATION: Secreted.
CC
CC
    -!- SIMILARITY: Belongs to the insulin family.
    _____
CC
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CC
    between the Swiss Institute of Bioinformatics and the EMBL outstation -
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    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
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    entities requires a license agreement (See http://www.isb-sib.ch/announce/
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    or send an email to license@isb-sib.ch).
    _____
CC
DR
    EMBL; X61092; CAA43405.1; -.
DR
    PIR; B42179; B42179.
    HSSP; P01308; 1AIO.
DR
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
DR
    Insulin family; Hormone; Glucose metabolism; Signal.
KW
FT
                      24
    SIGNAL
                1
                25
FT
    CHAIN
                      54
                              INSULIN B CHAIN.
              57
                     87
FT
    PROPEP
                             C PEPTIDE.
               90 110
FT
    CHAIN
                             INSULIN A CHAIN.
                     96
FT
    DISULFID
               31
                             INTERCHAIN.
               43 109
                             INTERCHAIN.
FT
    DISULFID
               95
                    100
FT
    DISULFID
SO
    SEQUENCE 110 AA; 12019 MW; 95A1F54BE7B247F9 CRC64;
 Query Match
                       46.5%; Score 273; DB 1; Length 110;
                       60.2%; Pred. No. 4.9e-22;
 Best Local Similarity
          53; Conservative 0; Mismatches 1; Indels 34; Gaps
          54 PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
            23 PAFVNQHLCGSHLVEALYLVCGERGFFYTPKTRREAEDPQVGQVELGGGPGAGSLQPLAL 82
Db
         86 -----RGIVEQCCTSICSLYQLENYCN 107
Qу
                  83 EGSLQKRGIVEQCCTSICSLYQLENYCN 110
```

```
RESULT 4
INS MACFA
ΙD
     INS MACFA
                   STANDARD;
                                  PRT;
                                          110 AA.
AC
     P30406; P01309;
     21-JUL-1986 (Rel. 01, Created)
DT
     13-AUG-1987 (Rel. 05, Last sequence update)
DT
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DE
     Insulin precursor.
GN
    INS.
    Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Primates; Catarrhini; Cercopithecidae;
OC
     Cercopithecinae; Macaca.
OC
OX
     NCBI TaxID=9541;
RN
     SEOUENCE FROM N.A.
RP
    MEDLINE=83080474; PubMed=6184262;
RX
RA
     Wetekam W., Groneberg J., Leineweber M., Wengenmayer F.,
RA
     Winnacker E.-L.;
RT
     "The nucleotide sequence of cDNA coding for preproinsulin from the
RT
     primate Macaca fascicularis.";
     Gene 19:179-183(1982).
RL
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
         increases cell permeability to monosaccharides, amino acids and
CC
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
         cycle, and glycogen synthesis in liver.
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
CC
        disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
     -!- SIMILARITY: Belongs to the insulin family.
CC
CC
     This SWISS-PROT entry is copyright. It is produced through a collaboration
CC
CC
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
CC
     the European Bioinformatics Institute. There are no restrictions on its
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     use by non-profit institutions as long as its content is in no way
     modified and this statement is not removed. Usage by and for commercial
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
CC
     ______
     EMBL; J00336; AAA36849.1; -.
DR
     PIR; JQ0178; JQ0178.
DR
DR
     HSSP; P01308; 1AI0.
     InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     Pfam; PF00049; Insulin; 1.
     PRINTS; PR00277; INSULINB.
DR
DR
     SMART; SM00078; IlGF; 1.
DR
     PROSITE; PS00262; INSULIN; 1.
KW
     Insulin family; Hormone; Glucose metabolism; Signal.
FT
     SIGNAL
                  1
                         24
FT
     CHAIN
                 25
                         54
                                  INSULIN B CHAIN.
FT
     PROPEP
                 57
                        87
                                 C PEPTIDE.
FT
     CHAIN
                  90
                       110
                                  INSULIN A CHAIN.
FT
     DISULFID
                  31
                       96
                                 INTERCHAIN.
FT
                  43
                       109
                                 INTERCHAIN.
     DISULFID
                  95
                       100
FT
     DISULFID
```

```
Query Match
                         46.5%; Score 273; DB 1; Length 110;
  Best Local Similarity 60.2%; Pred. No. 4.9e-22;
  Matches
           53; Conservative 0; Mismatches 1; Indels 34; Gaps
                                                                           1:
          54 PRFVNQHLCGSHLVEALYLVCGERGFFYTPKT----- 85
Qу
             23 PAFVNQHLCGSHLVEALYLVCGERGFFYTPKTRREAEDPOVGOVELGGGPGAGSLOPLAL 82
Db
          86 -----RGIVEQCCTSICSLYQLENYCN 107
Qу
                   11111111111111111111111111
          83 EGSLQKRGIVEQCCTSICSLYQLENYCN 110
Db
RESULT 5
INS ACOCA
    INS ACOCA
                   STANDARD;
                                 PRT;
                                          51 AA.
ID
AC
    P01324;
    21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
    Insulin.
GN
    INS.
    Acomys cahirinus (Egyptian spiny mouse).
OS
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Acomys.
OC
OX
    NCBI TaxID=10068;
RN
     [1]
RP
    COMPOSITION.
    MEDLINE=72189454; PubMed=5028210;
RX
RA
    Buenzli H.F., Humbel R.E.;
RT
     "Isolation and partial structural analysis of insulin from mouse (Mus
RT
    musculus) and spiny mouse (Acomys cahirinus).";
RL
    Hoppe-Seyler's Z. Physiol. Chem. 353:444-450(1972).
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
        increases cell permeability to monosaccharides, amino acids and
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
CC
    -!- SUBCELLULAR LOCATION: Secreted.
CC
    -!- SIMILARITY: Belongs to the insulin family.
DR
    PIR; A01591; INMSSP.
DR
    HSSP; P01308; 1TYM.
    InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     PRINTS; PR00277; INSULINB.
DR
     SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
KW
     Insulin family; Hormone; Glucose metabolism.
FT
     CHAIN
                 1
                        30
                                 INSULIN B CHAIN.
FT
    NON CONS
                 30
                        31
FT
    CHAIN
                 31
                        51
                                 INSULIN A CHAIN.
FT
    DISULFID
                 7
                        37
                                 INTERCHAIN (BY SIMILARITY).
FT
    DISULFID
                 19
                        50
                                 INTERCHAIN (BY SIMILARITY).
                       41
FT
    DISULFID
                 36
                                 BY SIMILARITY.
SQ
     SEQUENCE 51 AA; 5768 MW; 992BD8B629047D3D CRC64;
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SEQUENCE 110 AA; 11991 MW; 83C6E33A80A420F9 CRC64;

SO

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                                3; Mismatches
                                                  0; Indels
                                                                1; Gaps
                                                                            1;
Qу
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                                  PRT;
                                         110 AA.
AC
     P01308;
DT
     21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
     15-MAR-2004 (Rel. 43, Last annotation update)
DT
DE
    Insulin precursor.
GN
     INS.
OS
    Homo sapiens (Human).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
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RX
    MEDLINE=80120725; PubMed=6243748;
     Bell G.I., Pictet R.L., Rutter W.J., Cordell B., Tischer E.,
RA
    Goodman H.M.;
RA
     "Sequence of the human insulin gene.";
RT
RL
    Nature 284:26-32(1980).
RN
     [2]
RP
     SEQUENCE FROM N.A.
RX
    MEDLINE=80236313; PubMed=6248962;
    Ullrich A., Dull T.J., Gray A., Brosius J., Sures I.;
RA
RT
     "Genetic variation in the human insulin gene.";
RL
     Science 209:612-615(1980).
RN
     [3]
RP
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RX
    MEDLINE=80054779; PubMed=503234;
RA
     Bell G.I., Swain W.F., Pictet R.L., Cordell B., Goodman H.M.,
RA
    Rutter W.J.;
RT
     "Nucleotide sequence of a cDNA clone encoding human preproinsulin.";
RL
    Nature 282:525-527(1979).
RN
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RP
    SEQUENCE FROM N.A.
    MEDLINE=80147417; PubMed=6927840;
RX
RA
     Sures I., Goeddel D.V., Gray A., Ullrich A.;
RT
     "Nucleotide sequence of human preproinsulin complementary DNA.";
RL
    Science 208:57-59(1980).
RN
     [5]
RP
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RX
    MEDLINE=93364428; PubMed=8358440;
RA
    Lucassen A.M., Bell J.I., Julier C., Lathrop M.;
RT
     "Susceptibility to insulin dependent diabetes mellitus maps to a 4.1
RT
     kb segment of DNA spanning the insulin gene and associated VNTR.";
RL
    Nat. Genet. 4:305-310(1993).
RN
     [6]
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RP
     SEQUENCE FROM N.A.
RC
     TISSUE=Pancreas;
RX
     MEDLINE=22388257; PubMed=12477932;
     Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
RA
     Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,
RA
     Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
RA
     Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
RA
     Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,
RA
     Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,
RA
RA
     Brownstein M.J., Usdin T.B., Toshiyuki S., Carninci P., Prange C.,
RA
     Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullahy S.J.,
     Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,
RA
     Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,
RA
     Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,
RA
RA
     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
     Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,
RA
RA
     Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,
     Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,
RA
     Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smailus D.E.,
RA
     Schnerch A., Schein J.E., Jones S.J.M., Marra M.A.;
RA
     "Generation and initial analysis of more than 15,000 full-length
RT
RT
     human and mouse cDNA sequences.";
     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
RL
RN
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RP
RC
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     Fajardy I.I., Weill J.J., Stuckens C.C., Danze P.M.P.;
RA
     "Description of a novel RFLP diallelic polymorphism (-127 BsgI C/G)
RT
     within the 5' region of insulin gene.";
RT
RL
     Submitted (JUL-1998) to the EMBL/GenBank/DDBJ databases.
RN
     [8]
     SEQUENCE OF 25-54 AND 90-110.
RP
RA
     Nicol D.S.H.W., Smith L.F.;
RT
     "Amino-acid sequence of human insulin.";
RL
     Nature 187:483-485(1960).
RN
     SEQUENCE OF 57-87.
RP
     MEDLINE=71116410; PubMed=5101771;
RX
RA
     Oyer P.E., Cho S., Peterson J.D., Steiner D.F.;
RT
     "Studies on human proinsulin. Isolation and amino acid sequence of
RT
     the human pancreatic C-peptide.";
RL
     J. Biol. Chem. 246:1375-1386(1971).
RN
     [10]
RP
     SEQUENCE OF 57-87.
     MEDLINE=71257722; PubMed=5560404;
RX
     Ko A., Smyth D.G., Markussen J., Sundby F.;
RA
     "The amino acid sequence of the C-peptide of human proinsulin.";
RT
RL
     Eur. J. Biochem. 20:190-199(1971).
RN
     [11]
RP
     SYNTHESIS.
RX
     MEDLINE=75077277; PubMed=4443293;
RA
     Sieber P., Kamber B., Hartmann A., Joehl A., Riniker B., Rittel W.;
RT
     "Total synthesis of human insulin under directed formation of the
RT
     disulfide bonds.";
RL
     Helv. Chim. Acta 57:2617-2621(1974).
RN
     [12]
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SYNTHESIS OF 57-87.

RΡ

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RX
    MEDLINE=75040007; PubMed=4803504;
RA
    Naithani V.K.;
RT
     "Studies on polypeptides, IV. The synthesis of C-peptide of human
RT
    proinsulin.";
    Hoppe-Seyler's Z. Physiol. Chem. 354:659-672(1973).
RL
RN
     [13]
     SYNTHESIS OF 65-69 AND 70-73.
RP
    MEDLINE=73161263; PubMed=4698555;
RX
    Geiger R., Volk A.;
RA
RT
     "Synthesis of peptides with the properties of human proinsulin C
    peptides (hC peptide). 3. Synthesis of the sequences 14-17 and 9-13
RT
     of human proinsulin C peptides.";
RT
     Chem. Ber. 106:199-205(1973).
RL
RN
     [14]
RP
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    MEDLINE=73161261; PubMed=4698553;
RX
     Geiger R., Jaeger G., Keonig W., Treuth G.;
RA
RT
     "Synthesis of peptides with the properties of human proinsulin C
     peptides (hC peptide). I. Scheme for the synthesis and preparation of
RT
     the sequence 28-31 of human proinsulin C peptide.";
RT
RL
     Chem. Ber. 106:188-192(1973).
RN
     [15]
RP
     VARIANT LOS ANGELES SER-48.
    MEDLINE=84016053; PubMed=6312455;
RX
     Haneda M., Chan S.J., Kwok S.C.M., Rubenstein A.H., Steiner D.F.;
RA
     "Studies on mutant human insulin genes: identification and sequence
RT
     analysis of a gene encoding [SerB24]insulin.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 80:6366-6370(1983).
RL
RN
     [16]
     VARIANTS LOS ANGELES SER-48 AND CHICAGO LEU-49.
RΡ
    MEDLINE=84170233; PubMed=6424111;
RX
     Shoelson S., Fickova M., Haneda M., Nahum A., Musso G., Kaiser E.T.,
RA
     Rubenstein A.H., Tager H.;
RA
     "Identification of a mutant human insulin predicted to contain a
RT
     serine-for-phenylalanine substitution.";
RT
RL
     Proc. Natl. Acad. Sci. U.S.A. 80:7390-7394(1983).
RN
     [17]
RP
     VARIANT PROVIDENCE ASP-34.
RX
     MEDLINE=87175640; PubMed=3470784;
RA
     Chan S.J., Seino S., Gruppuso P.A., Schwartz R., Steiner D.F.;
RT
     "A mutation in the B chain coding region is associated with impaired
     proinsulin conversion in a family with hyperproinsulinemia.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 84:2194-2197 (1987).
RL
RN
     [18]
     VARIANT WAKAYAMA LEU-92.
RP
     MEDLINE=87058122; PubMed=3537011;
RX
     Sakura H., Iwamoto Y., Sakamoto Y., Kuzuya T., Hirata H.;
RA
RT
     "Structurally abnormal insulin in a diabetic patient. Characterization
RT
     of the mutant insulin A3 (Val-->Leu) isolated from the pancreas.";
RL
     J. Clin. Invest. 78:1666-1672(1986).
RN
     [19]
RP
     VARIANT HIS-89.
RX
     MEDLINE=90317021; PubMed=2196279;
     Barbetti F., Raben N., Kadowaki T., Cama A., Accili D., Gabbay K.H.,
RA
RA
     Merenich J.A., Taylor S.I., Roth J.;
     "Two unrelated patients with familial hyperproinsulinemia due to a
RT
     mutation substituting histidine for arginine at position 65 in the
RT
```

```
proinsulin molecule: identification of the mutation by direct
RT
    sequencing of genomic deoxyribonucleic acid amplified by polymerase
RT.
    chain reaction.";
    J. Clin. Endocrinol. Metab. 71:164-169(1990).
RL
RN
     [20]
RP
    VARIANT HIS-89.
    MEDLINE=85261996; PubMed=4019786;
RX
    Shibasaki Y., Kawakami T., Kanazawa Y., Akanuma Y., Takaku F.;
RA
     "Posttranslational cleavage of proinsulin is blocked by a point
RT
    mutation in familial hyperproinsulinemia.";
RT
    J. Clin. Invest. 76:378-380(1985).
RL
RN
     [21]
    VARIANT KYOTO LEU-89.
RP
    MEDLINE=92291307; PubMed=1601997;
RX
    Yano H., Kitano N., Morimoto M., Polonsky K.S., Imura H., Seino Y.;
RA
     "A novel point mutation in the human insulin gene giving rise to
RT
RT
    hyperproinsulinemia (proinsulin Kyoto).";
RL
     J. Clin. Invest. 89:1902-1907(1992).
RN
RP
     STRUCTURE BY NMR.
    MEDLINE=91104966; PubMed=2271664;
RX
     Hua Q.-X., Weiss M.A.;
RA
     "Toward the solution structure of human insulin: sequential 2D 1H NMR
RT
     assignment of a des-pentapeptide analogue and comparison with crystal
RT :
RT
     structure.";
     Biochemistry 29:10545-10555(1990).
RL
RN
     [23]
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RP
RX
     MEDLINE=91242467; PubMed=2036420;
RA
     Hua Q.-X., Weiss M.A.;
     "Comparative 2D NMR studies of human insulin and des-pentapeptide
RT
     insulin: sequential resonance assignment and implications for protein
RT
     dynamics and receptor recognition.";
RT
     Biochemistry 30:5505-5515(1991).
RL
RN
     [24]
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RP
     MEDLINE=91265527; PubMed=1646635;
RX
     Hua Q.-X., Weiss M.A.;
RA
     "Two-dimensional NMR studies of Des-(B26-B30)-insulin: sequence-
RT
     specific resonance assignments and effects of solvent composition.";
RT
     Biochim. Biophys. Acta 1078:101-110(1991).
RL
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Qу
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Qу
                  Db
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    01-APR-1993 (Rel. 25, Created)
DT
    01-APR-1993 (Rel. 25, Last sequence update)
DT
    15-MAR-2004 (Rel. 43, Last annotation update)
DT
DE
    Insulin precursor.
GN
    INS.
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OS
    Eukarvota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
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OC
    NCBI TaxID=9598;
OX
RN
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    SEOUENCE FROM N.A.
RP
    MEDLINE=92219953; PubMed=1560757;
RX
    Seino S., Bell G.I., Li W.;
RA
    "Sequences of primate insulin genes support the hypothesis of a
RT
    slower rate of molecular evolution in humans and apes than in
RT
RT
    monkeys.";
    Mol. Biol. Evol. 9:193-203(1992).
RL
RN
     [2]
    SEQUENCE FROM N.A.
RP
RX
    MEDLINE=22833521; PubMed=12952878;
    Stead J.D., Hurles M.E., Jeffreys A.J.;
RA
     "Global haplotype diversity in the human insulin gene region.";
RT
     Genome Res. 13:2101-2111(2003).
RL
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
        increases cell permeability to monosaccharides, amino acids and
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
CC
     -!- SIMILARITY: Belongs to the insulin family.
CC
     ______
CC
     This SWISS-PROT entry is copyright. It is produced through a collaboration
CC
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
CC
     the European Bioinformatics Institute. There are no restrictions on its
CC
     use by non-profit institutions as long as its content is in no way
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     modified and this statement is not removed. Usage by and for commercial
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
CC
     or send an email to license@isb-sib.ch).
CC
     ______
CC
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     EMBL; AY137497; AAN06933.1; -.
DR
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     PIR; A42179; A42179.
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     PDB; 1EFE; 29-MAR-00.
     InterPro; IPR004825; Ins/IGF/relax.
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DR
     PRINTS; PR00277; INSULINB.
DR
     SMART; SM00078; IlGF; 1.
DR
     PROSITE; PS00262; INSULIN; 1.
DR
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KW
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FT
     CHAIN
                 57
                       87
                                C PEPTIDE.
FT
     PROPEP
                 90 110
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FT
     CHAIN
                                INTERCHAIN.
FT
     DISULFID
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DISULFID 95
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                             INTERCHAIN.
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SO
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 Best Local Similarity 60.5%; Pred. No. 2.1e-21;
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                                            0; Indels 34; Gaps
 Matches
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Qу
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Db
         86 ----RGIVEQCCTSICSLYQLENYCN 107
Qу
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AC
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DT
    10-OCT-2003 (Rel. 42, Created)
    10-OCT-2003 (Rel. 42, Last sequence update)
DT
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
    Insulin precursor.
GN
    INS.
    Spermophilus tridecemlineatus (Thirteen-lined ground squirrel).
OS
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OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Sciuridae; Sciurinae;
OC
    Spermophilus.
OC
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OX
RN
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RP
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RC
    Tredrea M.M., Buck M.J., Guhaniyogi J., Squire T.L., Andrews M.T.;
RA
    "Regulation of PDK4 expression in a hibernating mammal.";
RT
    Submitted (JUN-2001) to the EMBL/GenBank/DDBJ databases.
RL
    -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
CC
        increases cell permeability to monosaccharides, amino acids and
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
    -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
CC
    -!- SUBCELLULAR LOCATION: Secreted.
CC
    -!- SIMILARITY: Belongs to the insulin family.
CC
    CC
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CC
    between the Swiss Institute of Bioinformatics and the EMBL outstation -
    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
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    modified and this statement is not removed. Usage by and for commercial
    entities requires a license agreement (See http://www.isb-sib.ch/announce/
CC
    or send an email to license@isb-sib.ch).
CC
    _____
CC
DR
    EMBL; AY038604; AAK72558.1; -.
    HSSP; P01308; 1LNP.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
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Pfam; PF00049; Insulin; 1.
DR
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
    PROSITE; PS00262; INSULIN; 1.
DR
    Insulin family; Hormone; Glucose metabolism; Signal.
KW
FT
    SIGNAL
                 1
                        24
                                BY SIMILARITY.
                 25
                        54
                                INSULIN B CHAIN.
FT
    CHAIN
                 57
                        87
                                C PEPTIDE.
    PROPEP
FT
                                INSULIN A CHAIN.
FT
                 90
                       110
    CHAIN
    DISULFID
                 31
                       96
                                INTERCHAIN (BY SIMILARITY).
FT
                                INTERCHAIN (BY SIMILARITY).
                 43
                       109
FT
    DISULFID
                     100
                                BY SIMILARITY.
                 95
FT
    DISULFID
     SEQUENCE 110 AA; 12004 MW; 4511768D6622BEE5 CRC64;
SO
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  Best Local Similarity
                         57.4%; Pred. No. 2.7e-21;
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Qу
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Db
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Qу
                         Db
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INS BALBO
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                                 PRT;
                                         51 AA.
ID
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AC
     21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DE
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GN
     INS.
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OS
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OC
     Mammalia; Eutheria; Cetartiodactyla; Cetacea; Mysticeti;
OC
     Balaenopteridae; Balaenoptera.
OC
     NCBI TaxID=9768;
OX
RN
     [1]
     SEOUENCE.
RP
     Ishihara Y., Saito T., Ito Y., Fujino M.;
RA
     "Structure of sperm- and sei-whale insulins and their breakdown by
RT
RT
     whale pepsin.";
     Nature 181:1468-1469(1958).
RL
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
         increases cell permeability to monosaccharides, amino acids and
CC
CC
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
         cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
         disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
CC
     -!- SIMILARITY: Belongs to the insulin family.
DR
     PIR; A01582; INWH1S.
     HSSP; P01317; 1APH.
DR
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InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     PRINTS; PR00277; INSULINB.
DR
     SMART; SM00078; IlGF; 1.
     PROSITE; PS00262; INSULIN; 1.
DR
     Insulin family; Hormone; Glucose metabolism.
KW
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FT
    CHAIN
                  1
    NON CONS
                  30
                         31
FT
                         51
    CHAIN
                  31
                                  INSULIN A CHAIN.
FT
                  7
                         37
                                  INTERCHAIN.
FT
     DISULFID
FT
                  19
                         50
                                  INTERCHAIN.
     DISULFID
FT
     DISULFID
                  36
                         41
                                  9007B50E400A7DDD CRC64;
SQ
     SEQUENCE
                51 AA;
                       5723 MW;
                          44.9%;
                                  Score 263.5; DB 1;
  Query Match
                                                      Length 51;
  Best Local Similarity
                          92.3%;
                                  Pred. No. 2.2e-21;
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           48; Conservative
                                 0; Mismatches
                                                   3; Indels
                                                                 1; Gaps
                                                                             1;
           56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
Qу
              1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASTCSLYQLENYCN 51
RESULT 10
INS CAMDR
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                    STANDARD;
                                   PRT;
                                           51 AA.
ID
     P01\overline{3}20;
AC
     21-JUL-1986 (Rel. 01, Created)
DT
     21-JUL-1986 (Rel. 01, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DΤ
DE
     Insulin.
GN
     INS.
     Camelus dromedarius (Dromedary) (Arabian camel).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Cetartiodactyla; Tylopoda; Camelidae; Camelus.
OC
OX
     NCBI TaxID=9838;
RN
     [1]
RP
     SEQUENCE.
     Danho W.O.;
RA
     "The isolation and characterization of insulin of camel (Camelus
RT
     dromedarius).";
RT
     J. Fac. Med. Baghdad 14:16-28(1972).
RL
CC
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
         increases cell permeability to monosaccharides, amino acids and
CC
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
         cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
         disulfide bonds.
     -!- SUBCELLULAR LOCATION: Secreted.
CC
CC
     -!- SIMILARITY: Belongs to the insulin family.
DR
     PIR; A92782; INCMA.
     HSSP; P01317; 2INS.
DR
     InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     PRINTS; PRO0277; INSULINB.
     SMART; SM00078; IlGF; 1.
DR
     PROSITE; PS00262; INSULIN; 1.
DR
     Insulin family; Hormone; Glucose metabolism.
KW
FT
     CHAIN
                   1
                         30
                                  INSULIN B CHAIN.
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FT
    NON CONS
                 30
                        31
FT
    CHAIN
                 31
                        51
                                 INSULIN A CHAIN.
FT
    DISULFID
                  7
                        37
                                 INTERCHAIN.
                 19
                        50
                                 INTERCHAIN.
FT
    DISULFID
FT
    DISULFID
                 36
                        41
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 Matches
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Qy
             1 FANQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASVCSLYQLENYCN 51
Db
RESULT 11
INS CAPHI
    INS CAPHI
                   STANDARD;
                                  PRT;
                                          51 AA.
ID
AC
    P01319;
    21-JUL-1986 (Rel. 01, Created)
DT
DT
    21-JUL-1986 (Rel. 01, Last sequence update)
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
    Insulin.
GN
    INS.
    Capra hircus (Goat).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Cetartiodactyla; Ruminantia; Pecora; Bovoidea;
OC
OC
    Bovidae; Caprinae; Capra.
    NCBI TaxID=9925;
OX
RN
    [1]
RP
    SEQUENCE.
RX
    MEDLINE=66160119; PubMed=5949593;
RA
    Smith L.F.;
    "Species variation in the amino acid sequence of insulin.";
RT
    Am. J. Med. 40:662-666(1966).
RL
    -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
CC
        increases cell permeability to monosaccharides, amino acids and
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
CC
        cycle, and glycogen synthesis in liver.
    -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
CC
CC
    -!- SUBCELLULAR LOCATION: Secreted.
    -!- SIMILARITY: Belongs to the insulin family.
CC
DR
    PIR; A01586; INGT.
DR
    HSSP; P01317; 1APH.
DR
    InterPro; IPR004825; Ins/IGF/relax.
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
    PROSITE; PS00262; INSULIN; 1.
DR
    Insulin family; Hormone; Glucose metabolism.
KW
                                 INSULIN B CHAIN.
FT
    CHAIN
                        30
                  1
FT
    NON CONS
                 30
                        31
                        51
                                 INSULIN A CHAIN.
FT
    CHAIN
                 31
                 7
                        37
FT
     DISULFID
                                 INTERCHAIN.
                 19
                        50
                                 INTERCHAIN.
FT
     DISULFID
FT
     DISULFID
                 36
                        41
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51 AA; 5692 MW; 9007B50CDB4E7DDD CRC64;
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                         44.9%; Score 263.5; DB 1; Length 51;
 Query Match
 Best Local Similarity 90.4%; Pred. No. 2.2e-21;
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                                                                1; Gaps
                                                                            1;
          56 FVNOHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
QУ
             1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCAGVCSLYQLENYCN 51
Db
RESULT 12
INS PIG
ΙD
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                   STANDARD;
                                  PRT;
                                         108 AA.
AC.
     P01315; Q9TSJ5;
     21-JUL-1986 (Rel. 01, Created)
     16-OCT-2001 (Rel. 40, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
     Insulin precursor.
GN
     INS.
     Sus scrofa (Pig).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Cetartiodactyla; Suina; Suidae; Sus.
OC
     NCBI TaxID=9823;
OX
RN
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RP
     Han X.G., Tuch B.E.;
RA
     "Complete porcine preproinsulin cDNA sequence.";
RT
     Submitted (MAY-1998) to the EMBL/GenBank/DDBJ databases.
RL
RN
     SEQUENCE FROM N.A.
RP
RC
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     MEDLINE=22135958; PubMed=12140686;
RX
     Amarger V., Nguyen M., Laere A.S., Braunschweig M., Nezer C.,
RA
     Georges M., Andersson L.;
RA
     "Comparative sequence analysis of the INS-IGF2-H19 gene cluster in
RT
RT
     pigs.";
RT.
     Mamm. Genome 13:388-398(2002).
RN
     [3]
     SEQUENCE OF 25-108.
RP
     MEDLINE=68286485; PubMed=5657063;
RX
RA
     Chance R.E., Ellis R.M., Bromer W.W.;
     "Porcine proinsulin: characterization and amino acid sequence.";
RT
RL
     Science 161:165-167(1968).
RN
     [4]
     REVISION TO 59.
RP
     Chance R.E.;
RA
     Submitted (JUL-1970) to the PIR data bank.
RL
RN
     X-RAY CRYSTALLOGRAPHY (1.9 ANGSTROMS).
RΡ
     Blundell T.L., Dodson G.G., Hodgkin D., Mercola D.;
RA
     "Insulin. The structure in the crystal and its reflection in
RT
     chemistry and biology.";
RT
     Adv. Protein Chem. 26:279-402(1972).
RL
RN
RP
     X-RAY CRYSTALLOGRAPHY (1.5 ANGSTROMS).
RA
     Isaacs N.W., Agarwal R.C.;
```

```
"Experience with fast Fourier least squares in the refinement of the
RT
RT
    crystal structure of rhombohedral 2-zinc insulin at 1.5-A
RT
     resolution.";
    Acta Crystallogr. A 34:782-791(1978).
RL
RN
RP
    X-RAY CRYSTALLOGRAPHY (1.5 ANGSTROMS).
    MEDLINE=89099318; PubMed=2905485;
RX
    Baker E.N., Blundell T.L., Cutfield J.F., Cutfield S.M., Dodson E.J.,
RA
    Dodson G.G., Crowfoot Hodgkin D.M., Hubbard R.E., Isaacs N.W.,
RA
    Reynolds C.D., Sakabe K., Sakabe N., Vijayan N.M.;
RA
     "The structure of 2Zn pig insulin crystals at 1.5-A resolution.";
RT
    Philos. Trans. R. Soc. Lond., B, Biol. Sci. 319:369-456(1988).
RL
RN
    X-RAY CRYSTALLOGRAPHY (2.0 ANGSTROMS).
RP
RX
    MEDLINE=92126280; PubMed=1772633;
    Balschmidt P., Hansen F.B., Dodson E., Dodson G., Korber F.;
RT
     "Structure of porcine insulin cocrystallized with clupeine Z.";
RL
    Acta Crystallogr. B 47:975-986(1991).
RN
RP
    X-RAY CRYSTALLOGRAPHY.
    MEDLINE=91222450; PubMed=2025410;
RX
    Badger J., Harris M.R., Reynolds C.D., Evans A.C., Dodson E.J.,
RA
     Dodson G.G., North A.C.T.;
RA
     "Structure of the pig insulin dimer in the cubic crystal.";
RT
    Acta Crystallogr. B 47:127-136(1991).
RL
RN
    [10]
    X-RAY CRYSTALLOGRAPHY (1.65 ANGSTROMS).
RP
     Diao J.-S., Wan Z.-L., Chang W.-R., Liang D.-C.;
RA
     "Structure of monomeric porcine DesB1-B2 despentapeptide (B26-B30)
RT
RT
     insulin at 1.65-A resolution.";
    Acta Crystallogr. D 53:507-512(1997).
RL
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
CC
         increases cell permeability to monosaccharides, amino acids and
         fatty acids. It accelerates glycolysis, the pentose phosphate
CC
CC
         cycle, and glycogen synthesis in liver.
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
CC
         disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
CC
     -!- SIMILARITY: Belongs to the insulin family.
CC
     -!- DATABASE: NAME=Protein Spotlight;
CC
        NOTE=Issue 9 of April 2001;
        WWW="http://www.expasy.org/spotlight/articles/sptlt009.html".
CC
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     the European Bioinformatics Institute. There are no restrictions on its
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
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     PDB; 3INS; 09-JAN-89.
DR
DR
     PDB; 4INS; 31-JUL-94.
DR
     PDB; 6INS; 31-JAN-94.
     PDB; 7INS; 31-JAN-94.
DR
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PDB; 9INS; 15-OCT-91.
DR
DR
    PDB; 1IZA; 15-OCT-91.
    PDB; 1IZB; 15-OCT-91.
DR
    PDB; 2TCI; 29-JAN-96.
DR
    PDB; 1MPJ; 29-JAN-96.
DR
    PDB; 3MTH; 29-JAN-96.
DR
    PDB; 1DEI; 16-JUN-97.
DR
    PDB; 1SDB; 01-APR-98.
DR
    PDB; 1WAV; 28-FEB-97.
DR
    PDB; 1ZEI; 16-FEB-99.
DR
    PDB; 1ZNI; 28-JAN-98.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
DR
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
DR
    PROSITE; PS00262; INSULIN; 1.
KW
    Insulin family; Hormone; Glucose metabolism; Signal; 3D-structure.
FT
    SIGNAL
                 1
                        24
FT
    CHAIN
                 25
                        54
                                INSULIN B CHAIN.
                 57
FT
    PROPEP
                        85
                                C PEPTIDE.
                 88
                       108
                                INSULIN A CHAIN.
FT
    CHAIN
    DISULFID
                 31
                      94
                                INTERCHAIN.
FT
                      107
                 43
                                INTERCHAIN.
FT
    DISULFID
                 93
                      98
FT
    DISULFID
                 26
                        46
FT
    HELIX
                 48
                        48
FT
    STRAND
FT
                 89
                        94
    HELIX
                100
FT
    HELIX
                       106
FT
    STRAND
                107
                      107
SQ
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Qу
             25 FVNQHLCGSHLVEALYLVCGERGFFYTPKARREAENPQAGAVELGGGLGGLQALALEGPP 84
          86 -- RGIVEQCCTSICSLYQLENYCN 107
Qу
               85 QKRGIVEQCCTSICSLYQLENYCN 108
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ID
    INS RABIT
                   STANDARD;
                                 PRT;
    P01311;
AC
DT
     21-JUL-1986 (Rel. 01, Created)
     01-FEB-1996 (Rel. 33, Last sequence update)
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
DΕ
    Insulin precursor.
GN
OS
    Oryctolagus cuniculus (Rabbit).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
    Mammalia; Eutheria; Lagomorpha; Leporidae; Oryctolagus.
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    NCBI TaxID=9986;
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RP
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    STRAIN=New Zealand white; TISSUE=Pancreas;
RC
    MEDLINE=94179230; PubMed=8132571;
RX
    Devaskar S.U., Giddings S.J., Rajakumar P.A., Carnaghi L.R.,
RA
RA
    Menon R.K., Zahm D.S.;
RT
    "Insulin gene expression and insulin synthesis in mammalian neuronal
RT
    cells.";
RL
    J. Biol. Chem. 269:8445-8454(1994).
RN
    SEQUENCE OF 25-54 AND 90-110.
RP
    MEDLINE=66160119; PubMed=5949593;
RX
RA
    Smith L.F.;
RT
    "Species variation in the amino acid sequence of insulin.";
RL
    Am. J. Med. 40:662-666(1966).
RN
RP
    SEQUENCE OF 56-110 FROM N.A.
RA
    Giddings S.J., Carnaghi L.R., Devaskar S.U.;
    Submitted (APR-1991) to the EMBL/GenBank/DDBJ databases.
RL
CC
    -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
        increases cell permeability to monosaccharides, amino acids and
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
    -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
    -!- SUBCELLULAR LOCATION: Secreted.
CC
CC
    -!- SIMILARITY: Belongs to the insulin family.
    _____
CC
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    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
    modified and this statement is not removed. Usage by and for commercial
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    entities requires a license agreement (See http://www.isb-sib.ch/announce/
    or send an email to license@isb-sib.ch).
CC
CC
    EMBL; U03610; AAA19033.1; -.
DR
DR
    EMBL; M61153; AAA17540.1; -.
    PIR; A53438; INRB.
DR
    HSSP; P01308; 1TYM.
DR
    InterPro; IPR004825; Ins/IGF/relax.
DR
    Pfam; PF00049; Insulin; 1.
DR
    PRINTS; PR00277; INSULINB.
    SMART; SM00078; IlGF; 1.
DR
DR
    PROSITE; PS00262; INSULIN; 1.
KW
    Insulin family; Hormone; Glucose metabolism; Signal.
FT
    SIGNAL
                 1
                        24
FT
    CHAIN
                 25
                        54
                                INSULIN B CHAIN.
    PROPEP
                 57
                       87
FT
                                C PEPTIDE.
FT
    CHAIN
                 90 110
                                INSULIN A CHAIN.
FT
    DISULFID
                31
                       96
                                INTERCHAIN.
FT
                43
                       109
                                INTERCHAIN.
    DISULFID
FT
                 95
    DISULFID
                       100
FT
                83
                      83
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SQ
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Best Local Similarity 59.3%; Pred. No. 5.6e-21;
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Qу
             25 FVNQHLCGSHLVEALYLVCGERGFFYTPKSRREVEELQVGQAELGGGPGAGGLQPSALEL 84
Db
          86 ----RGIVEQCCTSICSLYQLENYCN 107
QУ
                 Db
          85 ALQKRGIVEQCCTSICSLYQLENYCN 110
RESULT 14
INS FELCA
    INS FELCA
ID
                   STANDARD;
                                 PRT;
                                         51 AA.
AC
    P06306;
    01-JAN-1988 (Rel. 06, Created)
    01-JAN-1988 (Rel. 06, Last sequence update)
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
    Insulin.
GN
    INS.
OS
    Felis silvestris catus (Cat).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Carnivora; Fissipedia; Felidae; Felis.
OC
OX
    NCBI TaxID=9685;
RN
    [1]
    SEQUENCE.
RP
    MEDLINE=86214076; PubMed=3518635;
RX
    Hallden G., Gafvelin G., Mutt V., Joernvall H.;
RA
    "Characterization of cat insulin.";
RT
    Arch. Biochem. Biophys. 247:20-27(1986).
RL
CC
    -!- FUNCTION: Insulin decreases blood glucose concentration. It
        increases cell permeability to monosaccharides, amino acids and
CC
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
        disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
CC
    -!- SIMILARITY: Belongs to the insulin family.
DR
    PIR; A01588; INCT.
DR
    HSSP; P01317; 1APH.
    InterPro; IPR004825; Ins/IGF/relax.
    PRINTS; PR00277; INSULINB.
DR
    SMART; SM00078; IlGF; 1.
DR
DR
    PROSITE; PS00262; INSULIN; 1.
KW
    Insulin family; Hormone; Glucose metabolism.
                 1
FT
    CHAIN
                       30
                                INSULIN B CHAIN.
    NON CONS
                 30
                       31
FT
FT
    CHAIN
                 31
                       51
                                INSULIN A CHAIN.
                 7
                       37
FT
    DISULFID
                                INTERCHAIN.
                 19
                       50
FT
    DISULFID
                                INTERCHAIN.
FT
                 36
                       41
    DISULFID
               51 AA; 5745 MW; 9007B5096A0A7DDD CRC64;
SQ
     SEQUENCE
  Query Match
                        44.7%; Score 262.5; DB 1; Length 51;
  Best Local Similarity 90.4%; Pred. No. 2.8e-21;
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                                                              1; Gaps
                                                                         1;
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56 FVNQHLCGSHLVEALYLVCGERGFFYTPKTRGIVEQCCTSICSLYQLENYCN 107
QУ
             Db
           1 FVNQHLCGSHLVEALYLVCGERGFFYTPKA-GIVEQCCASVCSLYQLEHYCN 51
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INS CANFA
    INS CANFA
                  STANDARD;
                                 PRT: 110 AA.
AC
    P01321;
DT
    21-JUL-1986 (Rel. 01, Created)
DΤ
    21-JUL-1986 (Rel. 01, Last sequence update)
    10-OCT-2003 (Rel. 42, Last annotation update)
DE
    Insulin precursor.
GN
    INS.
    Canis familiaris (Dog).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Carnivora; Fissipedia; Canidae; Canis.
OC
OX
    NCBI TaxID=9615;
RN
    [1]
RP
    SEQUENCE FROM N.A.
    MEDLINE=83109071; PubMed=6296142;
RX
    Kwok S.C.M., Chan S.J., Steiner D.F.;
     "Cloning and nucleotide sequence analysis of the dog insulin gene.
RT
     Coded amino acid sequence of canine preproinsulin predicts an
RT
     additional C-peptide fragment.";
RT
     J. Biol. Chem. 258:2357-2363(1983).
RL
RN
     SEQUENCE OF 25-54 AND 90-110.
RP
    MEDLINE=66160119; PubMed=5949593;
RX
RA
     Smith L.F.:
     "Species variation in the amino acid sequence of insulin.";
RT
     Am. J. Med. 40:662-666(1966).
RL
     -!- FUNCTION: Insulin decreases blood glucose concentration. It
CC
        increases cell permeability to monosaccharides, amino acids and
CC
        fatty acids. It accelerates glycolysis, the pentose phosphate
CC
        cycle, and glycogen synthesis in liver.
CC
     -!- SUBUNIT: Heterodimer of a B chain and an A chain linked by two
CC
CC
        disulfide bonds.
CC
     -!- SUBCELLULAR LOCATION: Secreted.
     -!- SIMILARITY: Belongs to the insulin family.
CC
     _____
CC
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     or send an email to license@isb-sib.ch).
CC
CC
     EMBL; V00179; CAA23475.1; -.
DR
DR
     PIR; A92413; IPDG.
DR
     HSSP; P01317; 1APH.
     InterPro; IPR004825; Ins/IGF/relax.
DR
DR
     Pfam; PF00049; Insulin; 1.
     PRINTS: PRO0277; INSULINB.
DR
```

DR

SMART; SM00078; IlGF; 1.

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PROSITE; PS00262; INSULIN; 1.
    Insulin family; Hormone; Glucose metabolism; Signal.
KW
FT
                     24
    SIGNAL
               1
               25
                             INSULIN B CHAIN.
                     54
FT
    CHAIN
    PROPEP
               57
                    87
                             C PEPTIDE.
FT
               90
                             INSULIN A CHAIN.
                   110
FT
    CHAIN
               31
                    96
                             INTERCHAIN.
FT
    DISULFID
                   109
                             INTERCHAIN.
FT
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              43
              95
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FT
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Qу
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Db
         86 ----RGIVEQCCTSICSLYQLENYCN 107
Qу
               85 ALQKRGIVEQCCTSICSLYQLENYCN 110
Db
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